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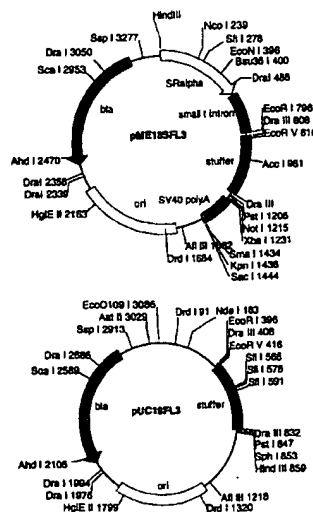
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(54) Primers for synthesizing full length cDNA clones and their use

(57) Primers for synthesizing full length cDNAs and their use are provided.

830 cDNA encoding a human protein has been isolated and nucleotide sequences of 5'-, and 3'-ends of the cDNA have been determined. Furthermore, primers for synthesizing the full length cDNA have been provided to clarify the function of the protein encoded by the cDNA. The full length cDNA of the present invention containing the translation start site provides information useful for analyzing the functions of the protein.

Figure 1



**Description****FIELD OF THE INVENTION**

- 5   **[0001]** The present invention relates to a polynucleotide encoding a novel protein, a protein encoded by the polynucleotide, and new uses of these.

**BACKGROUND OF THE INVENTION**

- 10   **[0002]** Currently, the sequencing projects, the determination and analysis of the genomic DNA of various living organisms have been in progress all over the world. The whole genomic sequences of more than 10 species of prokaryotes, a lower eukaryote, yeast, and a multicellular eukaryote, *C. elegans* are already determined. As to human genome, which is supposed to be composed of three thousand million base pairs, the world wide cooperative projects have been under way to analyze it, and the whole structure is predicted to be determined by the years 2002-2003. The aim  
15 of the determination of genomic sequence is to reveal the functions of all genes and their regulation and to understand living organisms as a network of interactions between genes, proteins, cells or individuals through deducing the information in a genome, which is a blueprint of the highly complicated living organisms. To understand living organisms by utilizing the genomic information from various species is not only important as an academic subject, but also socially significant from the viewpoint of industrial application.

- 20   **[0003]** However, determination of genomic sequences itself cannot identify the functions of all genes. For example, as for yeast, only the function of approximately half of the 6000 genes, which is predicted based on the genomic sequence, was able to be deduced. As for human, the number of the genes is predicted to be approximately one hundred thousand. Therefore, it is desirable to establish "a high throughput analysis system of the gene functions" which allows us to identify rapidly and efficiently the functions of vast amounts of the genes obtained by the genomic  
25 sequencing.

- [0004]** Many genes in the eukaryotic genome are split by introns into multiple exons. Thus, it is difficult to predict correctly the structure of encoded protein solely based on genomic information. In contrast, cDNA, which is produced from mRNA that lacks introns, encodes a protein as a single continuous amino acid sequence and allows us to identify the primary structure of the protein easily. In human cDNA research, to date, more than one million ESTs (Expression  
30 Sequence Tags) are publicly available, and the ESTs presumably cover not less than 80% of all human genes.

- [0005]** The information of ESTs is utilized for analyzing the structure of human genome, or for predicting the exon-regions of genomic sequences or their expression profile. However, many human ESTs have been derived from proximal regions to the 3'-end of cDNA, and information around the 5'-end of mRNA is extremely little. Among these human cDNAs, the number of the corresponding mRNAs whose encoding protein sequences are deduced is approximately  
35 7000, and further, the number of full-length therein is only 5500. Thus, even including cDNA registered as EST, the percentage of human cDNA obtained so far is estimated to be 10-15% of all the genes.

- [0006]** It is possible to identify the transcription start site of mRNA on the genomic sequence based on the 5'-end sequence of a full-length cDNA, and to analyze factors involved in the stability of mRNA that is contained in the cDNA, or in its regulation of expression at the translation stage. Also, since a full-length cDNA contains ATG, the translation  
40 start site, in the 5'-region, it can be translated into a protein in a correct frame. Therefore, it is possible to produce a large amount of the protein encoded by the cDNA or to analyze biological activity of the expressed protein by utilizing an appropriate expression system. Thus, analysis of a full-length cDNA provides valuable information which complements the information from genome sequencing. Also, full-length cDNA clones that can be expressed are extremely valuable in empirical analysis of gene function and in industrial application.

- 45   **[0007]** In particular, human secretory proteins or membrane proteins are would be useful by itself as a medicine like tissue plasminogen activator (TPA), or as a target of medicines like membrane receptors. In addition, genes for signal transduction-associated proteins (protein kinases, etc.), glycoprotein-associated proteins, transcription-associated proteins, and disease-associated proteins form a gene group rich in genes whose relationships to human diseases have been elucidated.

- 50   **[0008]** Therefore, it has great significance to isolate novel full-length cDNA clones of human, only few of which has been isolated. Especially, isolation of a novel cDNA clone encoding a secretory protein or membrane protein is desired since the protein itself would be useful as a medicine, and also the clones potentially include a gene associated with diseases. In addition, genes encoding proteins that are associated with signal transduction, glycoprotein, transcription, or diseases are expected to be useful as target molecules for therapy, or as medicines themselves. These genes form  
55 a gene group predicted to be strongly associated with diseases. Thus, identification of the full-length cDNA clones encoding those proteins has great significance.

SUMMARY OF THE INVENTION

**[0009]** An objective of the present invention is to provide a primer that enables synthesizing polynucleotide from human, the resulting polynucleotide or its clone, and a protein encoded by the polynucleotide.

**[0010]** The inventors have developed a method for efficiently cloning a human full-length cDNA that is predicted by the ATGpr etc. to be a full-length cDNA clone, from a full-length-enriched cDNA library that is synthesized by the oligo-capping method. Then, the inventors determined the nucleotide sequence of the obtained cDNA clones from both 5'- and 3'- ends. By utilizing the sequences, the inventors selected clones that were expected to contain a signal by the PSORT (Nakai K. and Kanehisa M. (1992) Genomics 14: 897-911), and obtained clones that contain a cDNA encoding a secretory protein or membrane protein. Moreover, the inventors specifically selected full-length cDNA clones that encode secretory or membrane proteins, signal transduction-associated proteins, glycoprotein-associated proteins, transcription-associated proteins, or disease-associated proteins from clones homologous to the clones in the Swiss-Prot ([http://www.ebi.ac.uk/ebi\\_docs/SwissProt\\_db/swisshome.html](http://www.ebi.ac.uk/ebi_docs/SwissProt_db/swisshome.html)) according to the keywords of SwissProt.

**[0011]** The full-length cDNA clones of the present invention have high fullness ratio since these were obtained by the combination of (1) construction of a full-length-enriched cDNA library that is synthesized by the oligo-capping method, and (2) a system in which fullness ratio is evaluated from the nucleotide sequence of the 5'-end (in this system, clones are selected based on the estimation by the ATGpr, following the removal of sequences judged not to be full-length when compared with ESTs). However, the primers of the present invention enable obtaining full-length cDNA easily without any special methods mentioned above.

**[0012]** Homology analysis in which the analysis is carried out against a non-full-length cDNA fragment to postulate the function of a protein encoded by said fragment, is being commonly performed. However, since such analysis is based on the information of the fragment, it is not clear as to whether this fragment corresponds to a part that is functionally important in the protein. In other words, the reliability of the homology analysis based on the information of a fragment is doubtful, as information relating to the structure of the whole protein is not available. However, the homology analysis of the present invention is conducted based on the information of a full-length cDNA comprising the whole coding region of the cDNA, and therefore, the homology of various portions of the protein can be analyzed. Hence, the reliability of the homology analysis has been dramatically improved in the present invention.

**[0013]** The inventors completed the invention by finding that it is possible to synthesize a novel full-length cDNA by using the combination of a primer that is designed based on the nucleotide sequence of the 5'-ends of the selected full-length cDNA clones and any of an oligo-dT primer or a 3'-primer that is designed based on the nucleotide sequence of the 3'-ends of the selected clones.

**[0014]** Thus, the present invention relates to primers described below, a method for synthesizing a polynucleotide using the primers, and polynucleotides obtained by the method.

**[0015]** First, the present invention relates to

(1) use of an oligonucleotide as a primer for synthesizing the polynucleotide comprising the nucleotide sequence set forth in any one of SEQ ID NOs: 1-829 and 2545, or the complementary strand thereof, wherein said oligonucleotide is complementary to said polynucleotide or the complementary strand thereof and comprises at least 15 nucleotides;

(2) a primer set for synthesizing polynucleotides, the primer set comprising an oligo-dT primer and an oligonucleotide complementary to the complementary strand of the polynucleotide comprising the nucleotide sequence set forth in any one of SEQ ID NOs: 1-829 and 2545, wherein said oligonucleotide comprises at least 15 nucleotides; and

(3) A primer set for synthesizing polynucleotides, the primer set comprising a combination of an oligonucleotide comprising a nucleotide sequence complementary to the complementary strand of the polynucleotide comprising a 5'-end nucleotide sequence and an oligonucleotide comprising a nucleotide sequence complementary to the polynucleotide comprising a 3'-end nucleotide sequence, wherein said oligonucleotides comprise at least 15 nucleotides and wherein said combination of 5'-end nucleotide sequence / 3'-end nucleotide sequence is selected from the combinations of 5'-end nucleotide sequence / 3'-end nucleotide sequence set forth in the SEQ ID NOs in Table 1.

**[0016]** Table 1 shows names of clones obtained in the examples described later, comprising the polynucleotide of the present invention (830 clones), names of nucleotide sequences at the 5'-end and 3'-end of the full-length cDNA, and their corresponding SEQ ID NOs. A blank indicates that the of the 3'-end sequence corresponding to the 5'-end sequence has not been determined the same clone.

**[0017]** The SEQ ID NO of a 5'-end sequence is shown on the right side of the name of the 5'-end sequence, and the SEQ ID NO of a 3'-end sequence is shown on the right side of the name of the 3'-end sequence.

Table 1

Correspondence between names of clone and the sequence name, and the SEQ ID.					
	Name of clone	Name of 5'-sequence	SEQ ID	3 Name of 3'-sequence	SEQ ID
5	BNGH41000020	F-BNGH41000020	1		
	BNGH41000087	F-BNGH41000087	2		
	BNGH41000091	F-BNGH41000091	3		
10	HEMBA1000006	F-HEMBA1000006	4	R-HEMBA1000006	830
	HEMBA1000121	F-HEMBA1000121	5	R-HEMBA1000121	831
	HEMBA1000128	F-HEMBA1000128	6	R-HEMBA1000128	832
	HEMBA1000275	F-HEMBA1000275	7	R-HEMBA1000275	833
	HEMBA1000300	F-HEMBA1000300	8	R-HEMBA1000300	834
15	HEMBA1000349	F-HEMBA1000349	9	R-nnnnnnnnnnnn	835
	HEMBA1000443	F-HEMBA1000443	10		
	HEMBA1000462	F-HEMBA1000462	11	R-HEMBA1000462	836
	HEMBA1000477	F-HEMBA1000477	12	R-HEMBA1000477	837
20	HEMBA1000590	F-HEMBA1000590	13	R-HEMBA1000590	838
	HEMBA1000634	F-HEMBA1000634	14	R-HEMBA1000634	839
	HEMBA1000671	F-HEMBA1000671	15	R-HEMBA1000671	840
	HEMBA1000713	F-HEMBA1000713	16	R-HEMBA1000713	841
	HEMBA1000732	F-HEMBA1000732	17	R-HEMBA1000732	842
25	HEMBA1000745	F-HEMBA1000745	18	R-nnnnnnnnnnnn	843
	HEMBA1000835	F-HEMBA1000835	19		
	HEMBA1000875	F-HEMBA1000875	20	R-HEMBA1000875	844
	HEMBA1000907	F-HEMBA1000907	21		
30	HEMBA1000940	F-HEMBA1000940	22	R-HEMBA1000940	845
	HEMBA1000962	F-HEMBA1000962	23	R-HEMBA1000962	846
	HEMBA1001184	F-HEMBA1001184	24	R-HEMBA1001184	847
	HEMBA1001221	F-HEMBA1001221	25	R-HEMBA1001221	848
	HEMBA1001228	F-HEMBA1001228	26	R-HEMBA1001228	849
35	HEMBA1001272	F-HEMBA1001272	27	R-HEMBA1001272	850
	HEMBA1001296	F-HEMBA1001296	28	R-HEMBA1001296	851
	HEMBA1001297	F-HEMBA1001297	29	R-HEMBA1001297	852
	HEMBA1001390	F-HEMBA1001390	30	R-HEMBA1001390	853
40	HEMBA1001563	F-HEMBA1001563	31	R-HEMBA1001563	854
	HEMBA1001621	F-HEMBA1001621	32	R-HEMBA1001621	855
	HEMBA1001878	F-HEMBA1001878	33	R-HEMBA1001878	856
	HEMBA1001886	F-HEMBA1001886	34	R-HEMBA1001886	857
	HEMBA1002048	F-HEMBA1002048	35	R-HEMBA1002048	858
45	HEMBA1002131	F-HEMBA1002131	36	R-HEMBA1002131	859
	HEMBA1002163	F-HEMBA1002163	37	R-HEMBA1002163	860
	HEMBA1002164	F-HEMBA1002164	38		
	HEMBA1002167	F-HEMBA1002167	39	R-HEMBA1002167	861
	HEMBA1002178	F-HEMBA1002178	40	R-HEMBA1002178	862
50	HEMBA1002195	F-HEMBA1002195	41	R-HEMBA1002195	863
	HEMBA1002227	F-HEMBA1002227	42	R-HEMBA1002227	864
	HEMBA1002239	F-HEMBA1002239	43		
	HEMBA1002316	F-HEMBA1002316	44	R-HEMBA1002316	865
55	HEMBA1002420	F-HEMBA1002420	45	R-HEMBA1002420	866
	HEMBA1002421	F-HEMBA1002421	46	R-HEMBA1002421	867
	HEMBA1002524	F-HEMBA1002524	47	R-HEMBA1002524	868



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Table 1 (continued)

Correspondence between names of clone and the sequence name, and the SEQ ID.					
	Name of clone	Name of 5'-sequence	SEQ ID	3 Name of 3'-sequence	SEQ ID
5	HEMBA1002551	F-HEMBA1002551	48	R-HEMBA1002551	869
	HEMBA1002767	F-HEMBA1002767	49	R-HEMBA1002767	870
	HEMBA1002985	F-HEMBA1002985	50	R-HEMBA1002985	871
	HEMBA1002992	F-HEMBA1002992	51		
10	HEMBA1003047	F-HEMBA1003047	52	R-HEMBA1003047	872
	HEMBA1003072	F-HEMBA1003072	53	R-HEMBA1003072	873
	HEMBA1003101	F-HEMBA1003101	54	R-HEMBA1003101	874
	HEMBA1003120	F-HEMBA1003120	55	R-HEMBA1003120	875
	HEMBA1003230	F-HEMBA1003230	56	R-HEMBA1003230	876
15	HEMBA1003294	F-HEMBA1003294	57	R-HEMBA1003294	877
	HEMBA1003315	F-HEMBA1003315	58	R-HEMBA1003315	878
	HEMBA1003392	F-HEMBA1003392	59	R-HEMBA1003392	879
	HEMBA1003399	F-HEMBA1003399	60	R-HEMBA1003399	880
20	HEMBA1003487	F-HEMBA1003487	61	R-HEMBA1003487	881
	HEMBA1003497	F-HEMBA1003497	62	R-HEMBA1003497	882
	HEMBA1003530	F-HEMBA1003530	63	R-HEMBA1003530	883
	HEMBA1003602	F-HEMBA1003602	64	R-HEMBA1003602	884
	HEMBA1003732	F-HEMBA1003732	65	R-HEMBA1003732	885
25	HEMBA1003945	F-HEMBA1003945	66	R-HEMBA1003945	886
	HEMBA1004007	F-HEMBA1004007	67	R-HEMBA1004007	887
	HEMBA1004067	F-HEMBA1004067	68		
	HEMBA1004085	F-HEMBA1004085	69	R-HEMBA1004085	888
30	HEMBA1004110	F-HEMBA1004110	70	R-nnnnnnnnnnnnn	889
	HEMBA1004250	F-HEMBA1004250	71	R-HEMBA1004250	890
	HEMBA1004391	F-HEMBA1004391	72	R-HEMBA1004391	891
	HEMBA1004444	F-HEMBA1004444	73	R-HEMBA1004444	892
	HEMBA1004454	F-HEMBA1004454	74	R-HEMBA1004454	893
35	HEMBA1004505	F-HEMBA1004505	75	R-HEMBA1004505	894
	HEMBA1004785	F-HEMBA1004785	76	R-HEMBA1004785	895
	HEMBA1004797	F-HEMBA1004797	77	R-HEMBA1004797	896
	HEMBA1004952	F-HEMBA1004952	78	R-HEMBA1004952	897
40	HEMBA1004971	F-HEMBA1004971	79	R-HEMBA1004971	898
	HEMBA1004982	F-HEMBA1004982	80	R-HEMBA1004982	899
	HEMBA1005070	F-HEMBA1005070	81	R-HEMBA1005070	900
	HEMBA1005084	F-HEMBA1005084	82	R-HEMBA1005084	901
	HEMBA1005145	F-HEMBA1005145	83	R-HEMBA1005145	902
45	HEMBA1005230	F-HEMBA1005230	84	R-HEMBA1005230	903
	HEMBA1005246	F-HEMBA1005246	85	R-HEMBA1005246	904
	HEMBA1005267	F-HEMBA1005267	86	R-HEMBA1005267	905
	HEMBA1005337	F-HEMBA1005337	87	R-HEMBA1005337	906
	HEMBA1005430	F-HEMBA1005430	88	R-HEMBA1005430	907
50	HEMBA1005449	F-HEMBA1005449	89	R-HEMBA1005449	908
	HEMBA1005489	F-HEMBA1005489	90	R-HEMBA1005489	909
	HEMBA1005522	F-HEMBA1005522	91	R-HEMBA1005522	910
	HEMBA1005545	F-HEMBA1005545	92	R-HEMBA1005545	911
55	HEMBA1005698	F-HEMBA1005698	93	R-HEMBA1005698	912
	HEMBA1005913	F-HEMBA1005913	94	R-HEMBA1005913	913
	HEMBA1005929	F-HEMBA1005929	95	R-HEMBA1005929	914

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Table 1 (continued)

Correspondence between names of clone and the sequence name, and the SEQ ID.					
	Name of clone	Name of 5'-sequence	SEQ ID	3 Name of 3'-sequence	SEQ ID
5	HEMBA1005945	F-HEMBA1005945	96	R-HEMBA1005945	915
	HEMBA1006016	F-HEMBA1006016	97	R-HEMBA1006016	916
	HEMBA1006171	F-HEMBA1006171	98	R-HEMBA1006171	917
	HEMBA1006276	F-HEMBA1006276	99	R-HEMBA1006276	918
10	HEMBA1006299	F-HEMBA1006299	100	R-HEMBA1006299	919
	HEMBA1006311	F-HEMBA1006311	101	R-HEMBA1006311	920
	HEMBA1006335	F-HEMBA1006335	102	R-HEMBA1006335	921
	HEMBA1006357	F-HEMBA1006357	103	R-HEMBA1006357	922
	HEMBA1006430	F-HEMBA1006430	104	R-HEMBA1006430	923
15	HEMBA1006482	F-HEMBA1006482	105	R-HEMBA1006482	924
	HEMBA1006517	F-HEMBA1006517	106	R-HEMBA1006517	925
	HEMBA1006544	F-HEMBA1006544	107	R-HEMBA1006544	926
	HEMBA1006572	F-HEMBA1006572	108	R-HEMBA1006572	927
20	HEMBA1006658	F-HEMBA1006658	109	R-HEMBA1006658	928
	HEMBA1006707	F-HEMBA1006707	110	R-HEMBA1006707	929
	HEMBA1006724	F-HEMBA1006724	111	R-HEMBA1006724	930
	HEMBA1006749	F-HEMBA1006749	112	R-HEMBA1006749	931
	HEMBA1006770	F-HEMBA1006770	113	R-HEMBA1006770	932
25	HEMBA1006902	F-HEMBA1006902	114	R-HEMBA1006902	933
	HEMBA1006912	F-HEMBA1006912	115	R-HEMBA1006912	934
	HEMBA1006916	F-HEMBA1006916	116	R-HEMBA1006916	935
	HEMBA1006960	F-HEMBA1006960	117	R-HEMBA1006960	936
30	HEMBA1007013	F-HEMBA1007013	118	R-HEMBA1007013	937
	HEMBA1007057	F-HEMBA1007057	119	R-HEMBA1007057	938
	HEMBA1007063	F-HEMBA1007063	120	R-HEMBA1007063	939
	HEMBA1007226	F-HEMBA1007226	121		
	HEMBA1007241	F-HEMBA1007241	122	R-HEMBA1007241	940
35	HEMBA1007291	F-HEMBA1007291	123	R-HEMBA1007291	941
	HEMBA1007332	F-HEMBA1007332	124	R-HEMBA1007332	942
	HEMBA1000106	F-HEMBA1000106	125	R-HEMBA1000106	943
	HEMBA1000276	F-HEMBA1000276	126	R-HEMBA1000276	944
40	HEMBA1000309	F-HEMBA1000309	127	R-HEMBA1000309	945
	HEMBA1000407	F-HEMBA1000407	128	R-HEMBA1000407	946
	HEMBA1000447	F-HEMBA1000447	129	R-HEMBA1000447	947
	HEMBA1000542	F-HEMBA1000542	130	R-HEMBA1000542	948
	HEMBA1000567	F-HEMBA1000567	131	R-HEMBA1000567	949
45	HEMBA1000642	F-HEMBA1000642	132	R-HEMBA1000642	950
	HEMBA1000668	F-HEMBA1000668	133	R-HEMBA1000668	951
	HEMBA1000679	F-HEMBA1000679	134	R-HEMBA1000679	952
	HEMBA1000881	F-HEMBA1000881	135	R-HEMBA1000881	953
	HEMBA1000905	F-HEMBA1000905	136	R-HEMBA1000905	954
50	HEMBA1001026	F-HEMBA1001026	137	R-HEMBA1001026	955
	HEMBA1001048	F-HEMBA1001048	138	R-HEMBA1001048	956
	HEMBA1001200	F-HEMBA1001200	139	R-HEMBA1001200	957
	HEMBA1001407	F-HEMBA1001407	140	R-HEMBA1001407	958
55	HEMBA1001530	F-HEMBA1001530	141	R-HEMBA1001530	959
	HEMBA1001547	F-HEMBA1001547	142	R-HEMBA1001547	960
	HEMBA1001573	F-HEMBA1001573	143	R-HEMBA1001573	961

Table 1 (continued)

Correspondence between names of clone and the sequence name, and the SEQ ID.				
Name of clone	Name of 5'-sequence	SEQ ID	3 Name of 3'-sequence	SEQ ID
Y79AA1001795	F-Y79AA1001795	816	R-Y79AA1001795	1560
Y79AA1001799	F-Y79AA1001799	817	R-Y79AA1001799	1561
Y79AA1001803	F-Y79AA1001803	818	R-Y79AA1001803	1562
Y79AA1001863	F-Y79AA1001863	819	R-Y79AA1001863	1563
Y79AA1002022	F-Y79AA1002022	820	R-Y79AA1002022	1564
Y79AA1002058	F-Y79AA1002058	821		
Y79AA1002121	F-Y79AA1002121	822	R-nnnnnnnnnnnnn	1565
Y79AA1002129	F-Y79AA1002129	823	R-nnnnnnnnnnnnn	1566
Y79AA1002213	F-Y79AA1002213	824	R-Y79AA1002213	1567
Y79AA1002334	F-Y79AA1002334	825	R-Y79AA1002334	1568
Y79AA1002373	F-Y79AA1002373	826	R-Y79AA1002373	1569
Y79AA1002376	F-Y79AA1002376	827	R-Y79AA1002376	1570
Y79AA1002378	F-Y79AA1002378	828	R-Y79AA1002378	1571
Y79AA1002381	F-Y79AA1002381	829	R-Y79AA1002381	1572
NT2RP2006580	F-NT2RP2006580	2545	R-NT2RP2006580	2546

The sequence name starting from "F" means the name of 5'-end sequence, and the sequence name starting from "R" means the name of 3'-end sequence. A blank indicates that the 3'-end sequence corresponding to the 5'-end sequence has not been determined in the clone.

[0018] Furthermore, the present invention relates to the use of the above primers, as described below.

(4) A polynucleotide which can be synthesized with the primer set of (2) or (3).

(5) A polynucleotide comprising a coding region in the polynucleotide of (4).

(6) A substantially pure protein encoded by polynucleotide of (4).

(7) A partial peptide of the protein of (6).

[0019] In addition, the present invention comprises a polynucleotide described below and a protein encoded by the polynucleotide.

(8) An isolated polynucleotide selected from the group consisting of

(a) a polynucleotide comprising a coding region of the nucleotide sequence set forth in any one of the SEQ ID NOs in Table 370;

(b) a polynucleotide comprising a nucleotide sequence encoding a protein comprising the amino acid sequence set forth in any one of the SEQ ID NOs in Table 370;

(c) a polynucleotide comprising a nucleotide sequence encoding a protein comprising an amino acid sequence selected from the amino acid sequences set forth in the SEQ ID NOs in Table 370, in which one or more amino acids are substituted, deleted, inserted, and/or added, wherein said protein is functionally equivalent to the protein comprising said amino acid sequence selected from the amino acid sequences set forth in the SEQ ID NOs in Table 370;

(d) a polynucleotide that hybridizes with a polynucleotide comprising a nucleotide sequence selected from the nucleotide sequences set forth in the SEQ ID NOs in Table 370, and that comprises a nucleotide sequence encoding a protein functionally equivalent to the protein encoded by the nucleotide sequence selected from the nucleotide sequences set forth in the SEQ ID NOs in Table 370;

(e) a polynucleotide comprising a nucleotide sequence encoding a partial amino acid sequence of a protein encoded by the polynucleotide of (a) to (d);

(f) a polynucleotide comprising a nucleotide sequence with at least 70% identity to the nucleotide sequence set forth in any one of the SEQ ID NOs in Table 370.

(9). A substantially pure protein encoded by the polynucleotide of (8).

(10) An antibody against the protein or peptide of any one of (6), (7), and (9).

(11) A vector comprising the polynucleotide of (5) or (8).

(12) A transformant carrying the polynucleotide of (5) or (8), or the vector of (11).

(13) A transformant expressively carrying the polynucleotide of (5) or (8), or the vector of (11).

(14) A method for producing the protein or peptide of any one of (6), (7), and (9), comprising culturing the transformant of (13) and recovering the expression product.

5 (15) An oligonucleotide comprising; the nucleotide sequence set forth in any one of the SEQ ID NOs in Table 370 or the nucleotide sequence complementary to the complementary strand thereof, wherein said oligonucleotide comprises 15 nucleotides or more.

(16) Use of the oligonucleotide of (15) as a primer for synthesizing a polynucleotide.

(17) Use of the oligonucleotide of (15) as a probe for detecting a gene.

10 (18) An antisense polynucleotide against the polynucleotide of (8), or the portion thereof.

(19) A method for synthesizing a polynucleotide, the method comprising:

a) synthesizing a complementary strand using a cDNA library as a template, and using the primer set of (2) or (3), or the primer of (16); and

15 b) recovering the synthesized product.

(20) The method of (19), wherein the cDNA library is obtainable by oligo-capping method.

(21) The method of (19), wherein the complementary strand is obtainable by PCR.

20 (22) A method for detecting the polynucleotide of (8), the method comprising:

a) incubating a target polynucleotide with the oligonucleotide of (15) under the conditions where hybridization occurs, and

b) detecting the hybridization of the target polynucleotide with the oligonucleotide of (15).

25 (23) A database of polynucleotides and/or proteins, the database comprising information on at least one sequence selected from the nucleotide sequences set forth in the SEQ ID NOs in Table 370 and/or the amino acid sequences set forth in the SEQ ID NOs in Table 370, or a medium on which the database is stored.

[0020] Any patents, patent applications, and publications cited herein are incorporated by reference.

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#### BRIEF DESCRIPTION OF THE DRAWINGS

[0021]

35 Figure 1 shows the restriction maps of vectors pME18SFL3 and pUC19FL3.

Figure 2 shows the reproducibility of gene expression analysis. The ordinate and the abscissa show the intensities of gene expression obtained in experiments different from each other.

Figure 3 shows the detection limit in gene expression analysis. The intensity of expression is shown in the ordinate, and the concentration ( $\mu\text{g/ml}$ ) of the probe used is shown in the abscissa.

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#### DETAILED DESCRIPTION OF THE INVENTION

[0022] Herein, "polynucleotide" is defined as a molecule in which multiple nucleotides are polymerized. There are no limitations in the number of the polymerized nucleotides. In case that the polymer contains relatively low number of nucleotides, it is also described as an "oligonucleotide". The polynucleotide or the oligonucleotide of the present invention can be a natural or chemically synthesized product. Alternatively, it can be synthesized using a template DNA by an enzymatic reaction such as PCR.

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[0023] All the cDNA provided by the invention are full-length cDNA. Herein, a "full-length cDNA" is defined as a cDNA which contains both ATG codon (the translation start site) and the stop codon. Accordingly, the untranslated regions, which are originally found in the upstream or downstream of the protein coding region in natural mRNA, may or may not be contained.

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An "isolated polynucleotide" is a polynucleotide the structure of which is not identical to that of any naturally occurring nucleic acid or to that of any fragment of a naturally occurring genomic nucleic acid spanning more than three separate genes. The term therefore covers, for example,

55

(a) a DNA which has the sequence of part of a naturally occurring genomic DNA molecule but is not flanked by both of the coding sequences that flank that part of the molecule in the genome of the organism in which it naturally occurs;

(b) a nucleic acid incorporated into a vector or into the genomic DNA of a prokaryote or eukaryote in a manner such that the resulting molecule is not identical to any naturally occurring vector or genomic DNA;

(c) a separate molecule such as a cDNA, a genomic fragment, a fragment produced by polymerase chain reaction (PCR), or a restriction fragment; and

(d) a recombinant nucleotide sequence that is part of a hybrid gene, i.e., a gene encoding a fusion protein. Specifically excluded from this definition are nucleic acids present in mixtures of different (i) DNA molecules, (ii) transfected cells, or (iii) cell clones: e.g., as these occur in a DNA library such as a cDNA or genomic DNA library.

**[0024]** The term "substantially pure" as used herein in reference to a given polypeptide means that the protein or polypeptide is substantially free from other biological macromolecules. The substantially pure protein or polypeptide is at least 75% (e.g., at least 80, 85, 95, or 99%) pure by dry weight. Purity can be measured by any appropriate standard method, for example, by column chromatography, polyacrylamide gel electrophoresis, or HPLC analysis.

**[0025]** All the clones of the present invention (830 clones) are novel and covering full-length, and also predicted to encode any of the following functional protein:

secretory proteins,  
membrane proteins,  
proteins associated to signal transduction (signal transduction-associated proteins; e.g. protein kinases, etc.),  
proteins associated to a glycoprotein (glycoprotein-associated proteins),  
proteins associated with transcription (transcription-associated proteins),  
proteins associated with diseases (disease-associated proteins),  
or, enzymes and/or metabolism-associated proteins, cell division- and/or cell proliferation-associated proteins,  
cytoskeleton-associated proteins, nuclear proteins, DNA-and/or RNA-binding proteins, ATP- and/or GTP-binding proteins, protein synthesis- and/or protein transport-associated proteins, and cellular defense-associated proteins.

**[0026]** Furthermore, all the cDNA clones of the present invention can be characterized as follows:

(1) a cDNA that is obtained by the oligo-capping method, which provides cDNA with high fullness ratio. The cDNA was selected by the score in the ATGpr (described as ATGpr1, as well), which is a program for prediction of the fullness of the 5'-end of cDNA based on the features of the 5'-end sequence. In addition, the PSORT, which is a program for prediction of the existence of the signal sequence selected, cDNA that contains a signal sequence in the 5'-end, or transmembrane region in the protein coding region. Furthermore, the homology search with the 5'-end sequences confirmed that, the selected clones were not identical to any of the known human mRNA (namely novel);

or,

(2) a cDNA that is obtained by the oligo-capping method, which provides cDNA with high fullness ratio. The cDNA was selected by the score in the ATGpr, which is a program for prediction of the fullness of the 5'-end based on the features of the 5'-end sequence. Furthermore, the a cDNA that has relative homology with an amino acid sequence of a protein with known functions was selected by the BLAST search (Altschul S.F., Gish W., Miller W., Myers E.W., and Lipman D.J. (1990) J. Mol. Biol. 215: 403-410 ; Gish W., and States D.J. (1993) Nature Genet. 3: 266-272) on the SwissProt database using the 5'-end sequence. In addition, the homology search using the 5'-end sequence confirmed that the selected clones were not identical to any of the known human mRNA (namely novel).

**[0027]** All clones are obtainable as a full-length clone by such a method as PCR (Current Protocols in Molecular Biology, Ausubel et al. edit, (1987) John Wiley & Sons, Section 6.1-6.4) using both the 5'- and 3'-end sequences, or using the 5'-end sequence and an oligo-dT primer that corresponds to the polyA sequence.

**[0028]** Specifically, PCR can be performed using an oligonucleotide that has 15 nucleotides longer, and specifically hybridizes with the complementary strand of the polynucleotide that contains the nucleotide sequence selected from the 5'-end sequences shown in Table 1 (SEQ ID NO: 1-829, and SEQ ID NO: 2545), and an oligo-dT primer as a 5'-, and 3'-primer, respectively. The length of the primers is usually 15-100 bp, and favorably between 15-35 bp. In case of LA PCR, which is described below, the primer length of 25-35 bp may provide a good result.

**[0029]** A method to design a primer that enables a specific amplification based on the given nucleotide sequence is known to those skilled in the art (Current Protocols in Molecular Biology, Ausubel et al. edit, (1987) John Wiley & Sons, Section 6.1-6.4). In designing a primer based on the 5'-end sequence, the primer is designed so as that, in principle, the amplification products will include the translation start site. Accordingly, in case that a given 5'-end nucleotide sequence is the 5'-untranslated region (5'UTR), any part of the sequence can be used as a 5'-primer as far as the specificity toward the target cDNA is insured. The translation start site can be predicted using a known method such

as the ATGpr as described below.

**[0030]** When synthesizing a polynucleotide, the target nucleotide sequence to be amplified can extend to several thousand bp in some cDNA. However, it is possible to amplify such a long nucleotides by using such as LA PCR (Long and Accurate PCR). It is advantageous to use LA PCR when synthesizing long DNA. In LA PCR, in which a special DNA polymerase having 3'  $\square$  5' exonuclease activity is used, misincorporated nucleotides can be removed. Accordingly, accurate synthesis of the complementary strand can be achieved even with a long nucleotide sequence. By using LA PCR, it is reported that amplification of a nucleotide with 20 kb longer can be achieved under desirable condition (Takeshi Hayashi (1996) Jikken-Igaku Bessatsu, "Advanced Technologies in PCR" Youdo-sha).

**[0031]** A template DNA for synthesizing the cDNA of the present invention can be obtained by using cDNA libraries that are prepared by various methods. The full-length cDNA clones obtained here are those with high fullness ratio, which were obtained using a combination of (1) a method to prepare a full-length-enriched cDNA library using the oligo-capping method, and (2) an estimation system for fullness using the 5'-end sequence (selection based on the estimation by the ATGpr after removing clones that are non-full-length compared to the ESTs). However, it is possible to easily obtain a full-length cDNA by using the primers that are provided by the present invention; not by the above described specialized method.

**[0032]** The problem with the cDNA libraries prepared by the known methods or commercially available is that mRNA contained in the libraries has very low fullness ratio. Thus, it is difficult to screen full-length cDNA clone directly from the library using ordinary cloning methods. The present invention has revealed a primer that is capable of synthesizing a full-length cDNA. If provided with primers, it is possible to synthesize a target full-length cDNA by using enzymatic reactions such as PCR. In particular, a full-length-enriched cDNA library, synthesized by methods such as oligo-capping, is desirable to synthesize a full-length cDNA with more reliability.

**[0033]** Once the nucleotide sequences of the full-length cDNAs obtained in the present invention is determined, it is possible to predict the functions of the proteins encoded by the cDNA clones, for example, by searching the databases such as GenBank (<http://www.ncbi.nlm.nih.gov/web/GenBank/>), Swiss-Prot ([http://www.ebi.ac.uk/ebi\\_docs/Swiss-Prot\\_db/swissprot.html](http://www.ebi.ac.uk/ebi_docs/Swiss-Prot_db/swissprot.html)), UniGene (<http://www.ncbi.nlm.nih.gov/UniGene/>) for homologies of the cDNAs, or by searching the amino acid sequences deduced from the full-length nucleotide sequences for signal sequence by using software such as PSORT (K. Nakai & M. Kanehisa, Genomics, 14: 897-991 (1992), for transmembrane region by using software such as SOSUI (T. Hirokawa et al., Bioinformatics, 14:378-379 (1998); Mitsui Knowledge Industry Co., Ltd.) or for motif by using software such as Pfam (<http://www.sanger.ac.uk/Software/Pfam/index.shtml>) or PROSITE (<http://www.expasy.ch/prosite>). As a matter of course, the functions are often predictable by using partial sequence information (preferably 300 nucleotides or more) instead of the full-length nucleotide sequences. However, the result of the prediction obtained by using partial sequence information does not always agree with the result obtained by using full-length nucleotide sequence, and thus it is needless to say that the prediction of function is preferably performed based on the full-length nucleotide sequences.

**[0034]** Homology search using each of GenBank, Swiss-Prot and UniGene was performed for the 826 clones whose full-length nucleotide sequences had been determined (HEMBA1005337, NT2RM1000407, NT2RM2001767, and NT2RP3003939 are not full-length). The amino acid sequences deduced from the full-length nucleotide sequences were searched for functional domains by using analytical software programs, PSORT, SOSUI and Pfam. Based on the results, proteins encoded by the cDNA clones were grouped into some categories and their functions were predicted.

**[0035]** The following 437 clones were categorized into secretory and/or membrane proteins. The clones categorized into secretory and/or membrane proteins are those which matched the full-length sequences of Swiss-Prot database with the keywords "growth factor", "cytokine", "hormone", "signal", "transmembrane", "membrane", "extracellular matrix", "receptor", "G-protein coupled receptor", "ionic channel", "voltage-gated channel", "calcium channel", "cell adhesion", "collagen" or "connective tissue"; those which matched the data, suggesting that the proteins are secretory and/or membrane proteins; or those which matched with the full-length sequences of GenBank or UniGene database similar description; and, further, those predicted to have an N-terminal signal sequence or a transmembrane region as a result of domain search for the amino acid sequences deduced from the full-length nucleotide sequences.

BNGH41000020, BNGH41000087, BNGH41000091, HEMBA1000121, HEMBA1000128, HEMBA1000349, HEMBA1000477, HEMBA1000590, HEMBA1000713, HEMBA1000732, HEMBA1000745, HEMBA1000835, HEMBA1000940, HEMBA1000962, HEMBA1001221, HEMBA1001228, HEMBA1001621, HEMBA1002131, HEMBA1002163, HEMBA1002167, HEMBA1002178, HEMBA1002195, HEMBA1002227, HEMBA1002420, HEMBA1002421, HEMBA1002767, HEMBA1003047, HEMBA1003101, HEMBA1003230, HEMBA1003392, HEMBA1003530, HEMBA1003602, HEMBA1003732, HEMBA1003945, HEMBA1004110, HEMBA1004250, HEMBA1004391, HEMBA1004444, HEMBA1004454, HEMBA1004505, HEMBA1004797, HEMBA1004982, HEMBA1005070, HEMBA1005449, HEMBA1005522, HEMBA1005545, HEMBA1005698, HEMBA1005945, HEMBA1006171, HEMBA1006299, HEMBA1006311, HEMBA1006335, HEMBA1006357, HEMBA1006430, HEMBA1006482, HEMBA1006707, HEMBA1006724, HEMBA1006749, HEMBA1006902, HEMBA1006960, HEMBA1007241, HEMBB1000407, HEMBB1000447, HEMBB1000567, HEMBB1000679, HEMBB1000881,

HEMBB1001026, HEMBB1001048, HEMBB1001407, HEMBB1001530, HEMBB1001573, HEMBB1001847,  
 HEMBB1001978, HEMBB1002041, HEMBB1002162, HEMBB1002245, HEMBB1002427, HEMBB1002693,  
 MAMMA1000102, MAMMA1000106, MAMMA1000118, MAMMA1000141, MAMMA1000204, MAMMA1000226,  
 MAMMA1000457, MAMMA1000473, MAMMA1000496, MAMMA1000591, MAMMA1000681, MAMMA1000810,  
 5 MAMMA1000986, MAMMA1000994, MAMMA1001043, MAMMA1001141, MAMMA1001237, MAMMA1001344,  
 MAMMA1001418, MAMMA1001893, MAMMA1001957, MAMMA1001978,  
 MAMMA1002070, MAMMA1002091, MAMMA1002095, MAMMA1002165, MAMMA1002234, MAMMA1002586,  
 MAMMA1002633, MAMMA1003126, NT2RM1000462, NT2RM1000542, NT2RM1000580, NT2RM1000855,  
 10 NT2RM1000858, NT2RM1000899, NT2RM2000241, NT2RM2000410, NT2RM2000423, NT2RM2000565,  
 NT2RM2001626, NT2RM2001792, NT2RM2001939, NT2RM2001941, NT2RM4000198, NT2RM4000284,  
 NT2RM4000417, NT2RM4000444, NT2RM4000587, NT2RM4000593, NT2RM4000648, NT2RM4000761,  
 NT2RM4000997, NT2RM4001325, NT2RM4001735, NT2RM4001768, NT2RM4001843, NT2RM4002352,  
 NT2RP1000050, NT2RP1000181, NT2RP1000261, NT2RP1000300, NT2RP1000325, NT2RP1000448,  
 NT2RP1000551, NT2RP1000613, NT2RP1000981, NT2RP1001004, NT2RP1001563, NT2RP2000479,  
 15 NT2RP2000533, NT2RP2000616, NT2RP2000649, NT2RP2000663, NT2RP2000694, NT2RP2000818,  
 NT2RP2000903, NT2RP2001200, NT2RP2001276, NT2RP2001480, NT2RP2001495, NT2RP2001514,  
 NT2RP2001755, NT2RP2001915, NT2RP2001956, NT2RP2002063, NT2RP2002188, NT2RP2002232,  
 NT2RP2002527, NT2RP2002533, NT2RP2002721, NT2RP2002824, NT2RP2002942, NT2RP2002976,  
 NT2RP2003042, NT2RP2003210, NT2RP2003383, NT2RP2003390, NT2RP2003469, NT2RP2003593,  
 20 NT2RP2003655, NT2RP2003664, NT2RP2003950, NT2RP2004179, NT2RP2004205, NT2RP2004495,  
 NT2RP2004524, NT2RP2004556, NT2RP2004606, NT2RP2004648, NT2RP2004794, NT2RP2005027,  
 NT2RP2005163, NT2RP2005181, NT2RP2005378, NT2RP2005463, NT2RP2005541, NT2RP2005597,  
 NT2RP2005666, NT2RP2005883, NT2RP2005994, NT2RP2006004,  
 NT2RP2006042, NT2RP2006269, NT2RP1006512, NT2RP2006580, NT2RP3000169, NT2RP3000171,  
 25 NT2RP3000304, NT2RP3000436, NT2RP3000460, NT2RP3000616, NT2RP3000676, NT2RP3000721,  
 NT2RP3000818, NT2RP3000907, NT2RP3000921, NT2RP3001012, NT2RP3001159, NT2RP3001195,  
 NT2RP3001240, NT2RP3001271, NT2RP3001322, NT2RP3001388, NT2RP3001560, NT2RP3001592,  
 NT2RP3001650, NT2RP3001738, NT2RP3001858, NT2RP3002015, NT2RP3002160, NT2RP3002311,  
 NT2RP3002342, NT2RP3002411, NT2RP3002737, NT2RP3002790, NT2RP3002836, NT2RP3002900,  
 30 NT2RP3002958, NT2RP3003000, NT2RP3003076, NT2RP3003354, NT2RP3003532, NT2RP3003535,  
 NT2RP3003614, NT2RP3004025, NT2RP3004075, NT2RP3004083, NT2RP3004130, NT2RP3004133,  
 NT2RP3004309, NT2RP3004345, NT2RP3004406, NT2RP3004481, NT2RP3004552, NT2RP3004625,  
 NT2RP3004647, NT2RP4001001, NT2RP4001009, NT2RP4001467, NT2RP4001879, NT2RP4002187,  
 NT2RP4002451, NT2RP4002750, OVARC1000003, OVARC1000105, OVARC1000298, OVARC1000307,  
 35 OVARC1000313, OVARC1000410, OVARC1000439, OVARC1000553, OVARC1000811, OVARC1000873,  
 OVARC1000956, OVARC1001030, OVARC1001163, OVARC1001336, OVARC1001570, OVARC1001607,  
 OVARC1001725, OVARC1001991, PLACE1000033, PLACE1000231, PLACE1000560, PLACE1000740,  
 PLACE1000912, PLACE1000914, PLACE1000927, PLACE1001016, PLACE1001123, PLACE1001183,  
 PLACE1001231, PLACE1001340, PLACE1001401, PLACE1001407, PLACE1001464, PLACE1001516,  
 40 PLACE1001536, PLACE1001564, PLACE1001655, PLACE1001795,  
 PLACE1001836, PLACE1001918, PLACE1001949, PLACE1002080, PLACE1002095, PLACE1002355,  
 PLACE1002374, PLACE1002518, PLACE1002547, PLACE1002726, PLACE1002905, PLACE1002911,  
 PLACE1002967, PLACE1003407, PLACE1003573, PLACE1003737, PLACE1003772, PLACE1003839,  
 PLACE1003845, PLACE1003852, PLACE1004279, PLACE1004282, PLACE1004441, PLACE1004450,  
 45 PLACE1004482, PLACE1004520, PLACE1004630, PLACE1004657, PLACE1004648, PLACE1004816,  
 PLACE1005003, PLACE1005005, PLACE1005031, PLACE1005383, PLACE1005410, PLACE1005426,  
 PLACE1005544, PLACE1005569, PLACE1005660, PLACE1005725, PLACE1005745, PLACE1005878,  
 PLACE1005927, PLACE1006071, PLACE1006093, PLACE1006208, PLACE1006277, PLACE1006290,  
 PLACE1006443, PLACE1006716, PLACE1006809, PLACE1006959, PLACE1007081, PLACE1007096,  
 50 PLACE1007296, PLACE1007626, PLACE1007845, PLACE1007881, PLACE1008359, PLACE1008469,  
 PLACE1008716, PLACE1008744, PLACE1008985, PLACE1009067, PLACE1009196, PLACE1009279,  
 PLACE1009527, PLACE1009546, PLACE1009600, PLACE1009982, PLACE1010011, PLACE1010078,  
 PLACE1010251, PLACE1010445, PLACE1010713, PLACE1010784, PLACE1010827, PLACE1010968,  
 PLACE1011116, PLACE1011181, PLACE1011236, PLACE1011516, PLACE1011708, PLACE3000181,  
 55 PLACE3000213, PLACE4000354, SKNMC1000004, SKNMC1000014, SKNMC1000082, THYRO1000036,  
 THYRO1000099, THYRO1000196, THYRO1000400, THYRO1000584, THYRO1000678, THYRO1000776,  
 THYRO1000795, THYRO1000956, THYRO1001102, THYRO1001113,  
 THYRO1001205, THYRO1001237, THYRO1001242, THYRO1001266, THYRO1001327, THYRO1001456,

THYRO1001478, THYRO1001523, THYRO1001529, THYRO1001641, THYRO1001702, THYRO1001725,  
 Y79AA1000207, Y79AA1000226, Y79AA1000270, Y79AA1000426, Y79AA1000521, Y79AA1000876,  
 Y79AA1000888, Y79AA1000959, Y79AA1001013, Y79AA1001212, Y79AA1001264, Y79AA1001328,  
 Y79AA1001426, Y79AA1001427, Y79AA1001430, Y79AA1001727, Y79AA1001787, Y79AA1001795,  
 5 Y79AA1001799, Y79AA1001803, Y79AA1002022, Y79AA1002058, Y79AA1002129, Y79AA1002213,  
 Y79AA1002373,

[0036] The following 146 clones were categorized into glycoprotein-associated proteins. The clones categorized into glycoprotein-associated proteins are those which matched the full-length sequences of Swiss-Prot database with the keyword "glycoprotein"; those which matched the data suggesting that the proteins are glycoprotein; or those which

10 matched the full-length sequences of GenBank or UniGene database with similar description.  
 BNGH41000087, BNGH41000091, HEMBA1000349, HEMBA1000590, HEMBA1000745, HEMBA1000835,  
 HEMBA1001221, HEMBA1001228, HEMBA1001621, HEMBA1002131, HEMBA1002178, HEMBA1002421,  
 HEMBA1002767, HEMBA1003230, HEMBA1003392, HEMBA1004250, HEMBA1004391, HEMBA1004444,  
 HEMBA1004505, HEMBA1005449, HEMBA1005522, HEMBA1005545, HEMBA1006707, HEMBA1006749,  
 15 HEMBA1006902, HEMBB1000679, HEMBB1000881, HEMBB1001048, HEMBB1002120, HEMBB1002245,  
 HEMBB1002427, MAMMA1000102, MAMMA1000591, MAMMA1000681, MAMMA1001043, MAMMA1001237,  
 MAMMA1002070, MAMMA1002586, MAMMA1003126, NT2RM1000462, NT2RM1000580, NT2RM2001792,  
 NT2RM2001818, NT2RM2001939, NT2RM2001941, NT2RM4000198, NT2RM4000284, NT2RM4000417,  
 NT2RM4000648, NT2RM4000997, NT2RM4001325, NT2RM4002352, NT2RP1000613, NT2RP1000981,  
 20 NT2RP1001004, NT2RP2000616, NT2RP2000694, NT2RP2000903, NT2RP2001480, NT2RP2001755,  
 NT2RP2002533, NT2RP2003042, NT2RP2003210, NT2RP2004205, NT2RP2004606, NT2RP2005027,  
 NT2RP2005181, NT2RP2005541, NT2RP2005597, NT2RP2005883, NT2RP2006004, NT2RP2006042,  
 NT2RP2006269, NT2RP3000304, NT2RP3000616, NT2RP3000921, NT2RP3001650, NT2RP3002160,  
 NT2RP3002737, NT2RP3002958, NT2RP3003000, NT2RP3003532, NT2RP3004130, NT2RP3004133,  
 25 NT2RP3004481, NT2RP3004552, NT2RP3004640, NT2RP4000108, NT2RP4001467, NT2RP4002750,  
 OVARC1000003, OVARC1000553, OVARC1000811, OVARC1000873, OVARC1001336, OVARC1001607,  
 OVARC1001991, PLACE1000033, PLACE1000740, PLACE1001016,  
 PLACE1001123, PLACE1001231, PLACE1001464, PLACE1001655, PLACE1001836, PLACE1002355,  
 PLACE1002374, PLACE1002905, PLACE1002911, PLACE1003573, PLACE1003737, PLACE1003772,  
 30 PLACE1003839, PLACE1004282, PLACE1004441, PLACE1004450, PLACE1004520, PLACE1004648,  
 PLACE1005003, PLACE1005426, PLACE1006071, PLACE1006073, PLACE1006290, PLACE1007081,  
 PLACE1007845, PLACE1008716, PLACE1008744, PLACE1008985, PLACE1010251, PLACE1010784,  
 PLACE1010968, PLACE1011116, PLACE3000181, PLACE3000213, PLACE4000354, THYRO1000036,  
 THYRO1000196, THYRO1000584, THYRO1000956, THYRO1001266, Y79AA1000270, Y79AA1000426,  
 35 Y79AA1001727, Y79AA1001795, Y79AA1002022, Y79AA1002213,

[0037] The following 57 clones were categorized into signal transduction-associated proteins. The clones categorized into signal transduction-associated proteins are those which matched the full-length sequences of Swiss-Prot database with the keywords "serine/threonine-protein kinase", "tyrosine-protein kinase" or "SH3 domain"; those which matched the data suggesting that the proteins are signal transduction-associated proteins (for example, "ADP-ribosylation factor"); or those which matched the full-length sequences of GenBank or UniGene database with similar description; and, further, those which was similarly predicted to be signal transduction-associated proteins based on the matching data of Pfam.

40 HEMBA1000006, HEMBA1002195, HEMBA1002227, HEMBA1002551, HEMBA1005084, HEMBA1005929,  
 HEMBA1006658, HEMBA1006916, MAMMA1000881, MAMMA1001150, MAMMA1001310, MAMMA1002142,  
 45 NT2RM2001902, NT2RP1001020, NT2RP1001031, NT2RP2001469, NT2RP2001529, NT2RP2001769,  
 NT2RP2003179, NT2RP2003545, NT2RP2004670, NT2RP3000011, NT2RP3000022, NT2RP3000172,  
 NT2RP3000201, NT2RP3000820, NT2RP3003527, NT2RP3003849, NT2RP3003874, NT2RP3004067,  
 NT2RP4000634, NT2RP4000962, OVARC1000255, OVARC1000529, OVARC1000916, OVARC1001338,  
 OVARC1001569, PLACE1002329, PLACE1003135, PLACE1003598, PLACE1005519, PLACE1006208,  
 50 PLACE1008282, PLACE1008297, PLACE1010081, PLACE1011364, PLACE1011824, THYRO1001457,  
 THYRO1001593, THYRO1001700, THYRO1001770, Y79AA1000777, Y79AA1000967, Y79AA1002376,  
 Y79AA1002381, HEMBB1000668, NT2RM4001377

[0038] The following 81 clones were categorized into transcription-associated proteins. The clones categorized into transcription-associated proteins are those which keywords "transcription regulation", "zinc finger" or "homeobox" matched the full-length sequences of Swiss-Prot database; those which the matched the data suggesting that the proteins were transcription-associated proteins; or those which matched the full-length sequences of GenBank or UniGene database with similar description; and, further, those which was similarly predicted to be transcription-associated proteins based on the matching data of Pfam.



HEMBA1000462, HEMBA1000671, HEMBA1001297, HEMBA1001390, HEMBA1001886, HEMBA1002048,  
 HEMBA1003120, HEMBA1003497, HEMBA1004785, HEMBA1005230, HEMBA1005246, HEMBA1006276,  
 HEMBA1006572, HEMBA1007226, HEMBB1000106, HEMBB1000905, HEMBB1001959, HEMBB1002051,  
 HEMBB1002661, MAMMA1001094, MAMMA1001532, MAMMA1001615, NT2RM1000789, NT2RM2000632,  
 5 NT2RM2000773, NT2RM4000326, NT2RP1000271, NT2RP1000468, NT2RP2000092, NT2RP2000610,  
 NT2RP2000712, NT2RP2000739, NT2RP2001538, NT2RP2001662, NT2RP2001817, NT2RP2001948,  
 NT2RP2002564, NT2RP2002974, NT2RP2003138, NT2RP2003302, NT2RP2003940, NT2RP2004108,  
 NT2RP2004847, NT2RP2005247, NT2RP2005391, NT2RP2005535, NT2RP2005774, NT2RP2005941,  
 NT2RP2006092, NT2RP3000148, NT2RP3000232, NT2RP3000378, NT2RP3000652, NT2RP3001976,  
 10 NT2RP3004090, NT2RP3004119, NT2RP3004294, OVARC1001049, OVARC1001086, OVARC1001132,  
 OVARC1001807, PLACE1000258, PLACE1000442, PLACE1000907, PLACE1003529, PLACE1004166,  
 PLACE1004168, PLACE1004887, PLACE1005250, PLACE1005682, PLACE1006079, PLACE1008549,  
 PLACE1011407, PLACE1011978, THYRO1000580, Y79AA1000030, Y79AA1001090, Y79AA1001523,  
 Y79AA1002334, Y79AA1002378, HEMBB1002302,

15 **[0039]** The following 85 clones were categorized into disease-associated proteins. The clones categorized into disease-associated proteins are those which matched the full-length sequences of Swiss-Prot database with the keywords "disease mutation" or "syndrome"; those which matched the data suggesting that the proteins are disease-associated proteins; or those which matched the full-length sequences of Swiss-Prot database and GenBank or UniGene database where the matched sequences are those of genes or proteins which had been deposited in the database of Online  
 20 Mendelian Inheritance in Man (OMIM) (<http://www.ncbi.nlm.nih.gov/Omim/>), which is a database of human genes and diseases.

BNGH41000020, HEMBA1000349, HEMBA1000590, HEMBA1000671, HEMBA1000835, HEMBA1001184,  
 HEMBA1001228, HEMBA1001886, HEMBA1003120, HEMBA1004250, HEMBA1005246, HEMBA1005267,  
 HEMBA1006707, HEMBA1006749, HEMBA1006902, HEMBA1006916, HEMBA1007013, HEMBB1002120,  
 25 MAMMA1000204, MAMMA1002080, NT2RM2000632, NT2RM2001126, NT2RM2001558, NT2RP1000271,  
 NT2RP1000465, NT2RP1000579, NT2RP2000447, NT2RP2000514, NT2RP2000739, NT2RP2001223,  
 NT2RP2001529, NT2RP2001562, NT2RP2002674, NT2RP2003369, NT2RP2004108, NT2RP2004205,  
 NT2RP2005535, NT2RP2005941, NT2RP2006004, NT2RP3000059, NT2RP3000125, NT2RP3000201,  
 NT2RP3000232, NT2RP3000616, NT2RP3000677, NT2RP3000838, NT2RP3000921, NT2RP3001542,  
 30 NT2RP3002286, NT2RP3002721, NT2RP3002737, NT2RP3002738, NT2RP3004481, OVARC1000208,  
 OVARC1000275, OVARC1000331, OVARC1000410, OVARC1001086, OVARC1001132, OVARC1001607,  
 OVARC1001725, OVARC1001952, PLACE1000258, PLACE1000442, PLACE1000907, PLACE1001100,  
 PLACE1001500, PLACE1002905, PLACE1002967, PLACE1003407, PLACE1003428, PLACE1005005,  
 PLACE1005239, PLACE1005815, PLACE1007028, PLACE1008716, PLACE1011407, PLACE1011978,  
 35 PLACE2000118, THYPO1000580, THYRO1000866, THYRO1001071, THYRO1001478, Y79AA1001062,  
 Y79AA1001530,

**[0040]** It is unclear, by the analyses so far, whether or not the remaining 212 clones encode proteins belonging to any of the categories of secretory and/or membrane proteins, glycoprotein-associated proteins, signal transduction-associated proteins, transcription-associated proteins or disease-associated proteins. Nonetheless, it is still possible  
 40 for these clones to encode secretory and/or membrane proteins, glycoprotein-associated proteins, signal transduction-associated proteins, transcription-associated proteins, or disease-associated proteins. On the other hand, some of these clones can be presumed to have functions other than those as secretory and/or membrane proteins, glycoprotein-associated proteins, signal transduction-associated proteins, transcription-associated proteins and disease-associated proteins.

45 **[0041]** Among the 212 clones, the following clones presumably belong to the categories of enzymes and/or metabolism-associated proteins, cell division- and/or cell proliferation-associated proteins, cytoskeleton-associated proteins, nuclear proteins, DNA- and/or RNA-binding proteins, ATP-and/or GTP-binding proteins, protein synthesis- and/or protein transport-associated proteins, or cellular defense-associated proteins, although it is unclear whether or not the clones belong to any of the categories of secretory and/or transmembrane proteins, glycoprotein-associated proteins,  
 50 signal transduction-associated proteins, transcription-associated proteins, and disease-associated proteins.

**[0042]** The following 10 clones presumably belong to the category of enzymes and/or metabolism-associated proteins. The clones herein defined as clones presumably belonging to the category of enzymes and/or metabolism-associated proteins matched data containing keywords such as "metabolism", "oxidoreductase" and "E.C. No. (Enzyme commission number)".

55 HEMBA1003315, HEMBB1002465, MAMMA1000614, NT2RP2000178, NT2RP2001388, NT2RP2001903,  
 NT2RP2002304, NT2RP2005878, NT2RP3001685, PLACE1006219

**[0043]** The following 4 clones presumably belong to the category of cell division- and/or cell proliferation-associated proteins. The cDNA clones were herein defined as clones presumably belonging to the category of cell division- and/

or cell proliferation-associated proteins matched data containing keywords such as "cell division", "cell cycle", "mitosis", "chromosomal protein", "cell growth" and "apoptosis".

MAMMA1000403, NT2RM2000497, NT2RP2000394, Y79AA1002121

5 [0044] The following 6 clones presumably belong to the category of cytoskeleton-associated proteins. The cDNA clones were herein defined as clones presumably belonging to the category of cytoskeleton-associated proteins matched data containing keywords such as "structural protein", "cytoskeleton", "actin-binding" and "microtubules".

MAMMA 1001609, NT2RM2000589, NT2RP3000063, PLACE 1004078, PLACE 1004492, PLACE 1008657

10 [0045] The following 7 clones presumably belong to the category of nuclear proteins. The cDNA clones were herein defined as clones presumably belonging to the category of nuclear proteins matched data containing keywords such as "nuclear protein".

HEMBA1001878, HEMBA1002992, MAMMA1000614, NT2RM4000965, NT2RM2001738, NT2RP2001388, Y79AA1002121

15 [0046] The following 5 clones presumably belong to the category of DNA- and/or RNA-binding proteins. The cDNA clones were herein defined as clones presumably belonging to the category of DNA- and/or RNA-binding proteins matched data containing keywords such as "DNA-binding" and "RNA-binding".

HEMBA1003072, HEMBA1006770, HEMBA1007332, NT2RM2000497, Y79AA1002121

[0047] The following 7 clones presumably belong to the category of ATP- and/or GTP-binding proteins. The cDNA clones were herein defined as clones presumably belonging to the category of ATP- and/or GTP-binding proteins matched data containing keywords such as "ATP-binding" and "GTP-binding".

20 HEMBA1002316, MAMMA1001609, NT2RM2000306, NT2RM2000497, NT2RP2000178, NT2RP3003729, PLACE1004305

25 [0048] The following 7 clones presumably belong to the category of protein synthesis- and/or protein transport-associated proteins. The cDNA clones were herein defined as clones presumably belonging to the category of protein synthesis-associated and/or protein transport-associated proteins matched data containing keywords such as "translation regulation", "protein biosynthesis", "amino-acid biosynthesis", "ribosomal protein", "protein transport" and "signal recognition particle".

NT2RM4000965, NT2RP2005069, NT2RP3000481, NT2RP3000789, NT2RP4001877, OVARC1001833, OVARC1002058,

30 [0049] The following 1 clone presumably belongs to the category of cellular defense-associated proteins. The cDNA clones were herein defined as clones presumably belonging to the category of cellular defense-associated proteins matched data containing keywords such as "heat shock", "DNA repair" and "DNA damage".

PLACE1005539

35 [0050] Although it is unclear whether or not 26 out of 174 clones other than the above-mentioned clones belong to any of the above-described categories, these clones are predicted to have some functions, based on the homology search using their full-length sequences thereof. The clone names and the gene definitions found in the result of homology search are shown below, separated with a double-slash mark, //.

40 HEMBA1000634//Homo sapiens T-cell activation protein (PGR1) gene, complete cds.  
HEMBA1002524//Human MHC Class I region proline rich protein mRNA, complete cds.  
HEMBA1003399//MVP1 PROTEIN.  
HEMBA1005489//Mus musculus semaphorin cytoplasmic domain-associated protein 3A (Semcap3) mRNA, complete cds.  
HEMBA1000542//Mus musculus bromodomain-containing protein BP75 mRNA, complete cds.  
MAMMA1000788//Bos taurus P14 (p14) mRNA, complete cds.  
45 MAMMA1002128//ABC1 PROTEIN HOMOLOG PRECURSOR.  
NT2RM2000514//Homo Sapiens F-box protein Fbx21 (FBX21) mRNA, complete cds.  
NT2RM2000622//Mus musculus F-box protein FBL10 mRNA, partial cds.  
NT2RM4000100//Homo sapiens Leman coiled-coil protein (LCCP) mRNA, complete cds.  
NT2RP2005425//Homo sapiens mRNA for AKAP450 protein.  
50 NT2RP3001170//Mus musculus activity-dependent neuroprotective protein (Adnp) mRNA, complete cds.  
NT2RP3002571//Bos taurus mRNA for lyncein.  
NT2RP3004557//Human Ki nuclear autoantigen mRNA, complete cds.  
OVARC1001596//Homo sapiens Arf-like 2 binding protein BART1 mRNA, complete cds.  
PLACE1002153//Homo sapiens TACC2 protein (TACC2) mRNA, partial cds.  
55 PLACE1003163//Homo sapiens DBI-related protein mRNA, complete cds.  
PLACE1005736//Human mRNA for BAS-GRIP protein.  
PLACE1007702//Mus musculus TRA1 mRNA, complete cds.  
PLACE1011045//Homo sapiens E1-like protein mRNA, complete cds.

THYRO1000061//Mus musculus mRNA for UBE-1c1, UBE-1c2, UBE-1c3, complete cds.

THYRO1000964//Drosophila melanogaster Pelle associated protein Pellino (Pli) mRNA, complete cds.

Y79AA1000776//Mus musculus mRNA for GSG1, complete cds.

Y79AA1001056//Homo sapiens MAID protein mRNA, complete cds.

5 Y79AA1001272//Homo sapiens retinoic acid repressible protein (RARG-1) mRNA, complete cds.

Y79AA1001793//Mus musculus mRNA for GSG1, complete cds.

**[0051]** So far, useful information for presuming the functions are unavailable for the remaining 148 clones, whose names are listed below.

10 HEMBA1000275, HEMBA1000300, HEMBA1000443, HEMBA1000875, HEMBA1000907, HEMBA1001272,  
HEMBA1001296, HEMBA1001563, HEMBA1002164, HEMBA1002239, HEMBA1002985, HEMBA1003294,  
HEMBA1003487, HEMBA1004007, HEMBA1004067, HEMBA1004085, HEMBA1004952, HEMBA1004971,  
HEMBA1005145, HEMBA1005430, HEMBA1005913, HEMBA1006016, HEMBA1006517, HEMBA1006544,  
HEMBA1006912, HEMBA1007057, HEMBA1007063, HEMBA1007291, HEMBB1000276, HEMBB1000309,  
15 HEMBB1000642, HEMBB1001200, HEMBB1001547, HEMBB1002039, HEMBB1002228, HEMBB1002663,  
MAMMA1000046, MAMMA1000449, MAMMA1000528, MAMMA1000652, MAMMA1000706, MAMMA1000814,  
MAMMA1001066, MAMMA1001284, MAMMA1001623, MAMMA1001634, MAMMA1001901, MAMMA1002087,  
MAMMA1002205, MAMMA1002224, NT2RM2000582, NT2RM2001643, NT2RM4000115, NT2RM4000295,  
NT2RM4001321, NT2RP1000002, NT2RP1000239, NT2RP1000679, NT2RP1000740, NT2RP1000903,  
20 NT2RP2000240, NT2RP2001878, NT2RP2001921, NT2RP2002015, NT2RP2002409, NT2RP2002510,  
NT2RP2003599, NT2RP2003931, NT2RP2004069, NT2RP2004141, NT2RP2004447, NT2RP2004837,  
NT2RP2005514, NT2RP2005632, NT2RP2005887, NT2RP2006099, NT2RP2006134, NT2RP3000427,  
NT2RP3000444, NT2RP3000645, NT2RP3000871, NT2RP3001044, NT2RP3001061, NT2RP3001754,  
NT2RP3002281, NT2RP3002324, NT2RP3002353, NT2RP3002409, NT2RP3002448, NT2RP3002664,  
25 NT2RP3002887, NT2RP3002983, NT2RP3003448, NT2RP3003469, NT2RP3003473, NT2RP3003559,  
NT2RP3003963, NT2RP3004000, NT2RP3004202, NT2RP3004321,  
NT2RP3004355, NT2RP3004374, NT2RP4002715, OVARC1000090, OVARC1000137, OVARC1000467,  
OVARC1000775, OVARC1000853, OVARC1000995, OVARC1001222, OVARC1001260, OVARC1001727,  
OVARC1002178, PLACE1000986, PLACE1001114, PLACE1001229, PLACE1001788, PLACE1003438,  
30 PLACE1003460, PLACE1003644, PLACE1004028, PLACE1004199, PLACE1004519, PLACE1005601,  
PLACE1005669, PLACE1005768, PLACE1006515, PLACE1006786, PLACE1007040, PLACE1007077,  
PLACE1007591, PLACE1007971, PLACE1008984, PLACE1009735, PLACE2000219, PLACE4000455,  
THYRO1000846, THYRO1000999, THYRO1001063, THYRO1001128, THYRO1001471, THYRO1001495,  
THYRO1001608, THYRO1001803, Y79AA1000127, Y79AA1000750, Y79AA1001592, Y79AA1001863,

35 **[0052]** In the 437 clones categorized into secretory and/or membrane proteins by using their full-length sequences, 410 clones were also predicted to encode proteins having functions of secretory and/or membrane proteins by using their partial nucleotide sequences. In the 146 clones categorized into glycoprotein-associated proteins by using their full-length sequences, 124 clones were also predicted to encode proteins having functions of glycoprotein-associated proteins by using their partial nucleotide sequences. In the 57 clones categorized into signal transduction-associated proteins by using their full-length sequences, 46 clones were also predicted to encode proteins having functions of signal transduction-associated proteins by using their partial nucleotide sequences. In the 81 clones categorized into transcription-associated proteins by using their full-length sequences, 57 clones were also predicted to encode proteins having functions of transcription-associated proteins by using their partial nucleotide sequences. In the 85 clones categorized into disease-associated proteins by using their full-length sequences, 6 clones were also predicted to encode proteins having functions of disease-associated proteins by using their partial nucleotide sequences. The number of clones, which were predicted to encode disease-associated proteins based on the full-length nucleotide sequences, is much greater than that predicted based on the partial sequences. The reason is that the full-length sequences were categorized by using the data found in the OMIM database into the category of disease-associated proteins.

50 **[0053]** In some cases, the predicted functions based on the partial sequences are different from those based on the full-length sequences. The reason is that a protein does not always belong solely to a single category of the above-described functional categories, and therefore, it is possible for the protein to belong to both of the predicted functional categories. Besides, additional functions can be found for the clones classified into these functional categories by further analyses.

55 **[0054]** The following list shows the cDNA clones predicted and selected on the basis of the partial sequences (5' sequences) as cDNAs encoding secretory and/or membrane proteins, glycoprotein-associated proteins, signal transduction-associated proteins, transcription-associated proteins, or disease-associated proteins.

**[0055]** The clones that are selected by the score in the ATGpr and by the PSORT for the existence of a signal sequence can be expected to encode a secretory or membrane protein since they are predicted to possess the secretion

signal or a transmembrane region. The clones that are selected by the score in the ATGpr and by the PSORT for the existence of a signal sequence are listed below (254 clones).

5 HEMBA1000300 HEMBA1000713 HEMBA1000907  
 HEMBA1000962 HEMBA1001272 HEMBA1001297  
 HEMBA1002164 HEMBA1002239 HEMBA1002420  
 HEMBA1002421 HEMBA1003101 HEMBA1003294  
 HEMBA1003399 HEMBA1003602 HEMBA1003732  
 10 HEMBA1004110 HEMBA1004797 HEMBA1005430  
 HEMBA1006016 HEMBA1006171 HEMBA1006311  
 HEMBA1006335 HEMBA1006357 HEMBA1006572  
 HEMBA1006658 HEMBA1006707 HEMBA1006902  
 HEMBA1006960 HEMBA1007013 HEMBB1000276  
 HEMBB1000447 HEMBB1000567 HEMBB1000642  
 15 HEMBB1000905 HEMBB1001200 HEMBB1001407  
 HEMBB1001530 HEMBB1001547 HEMBB1001978  
 HEMBB1002162 HEMBB1002228 HEMBB1002245  
 HEMBB1002427 HEMBB1002465 HEMBB1002663  
 HEMBB1002693 MAMMA1000046 MAMMA1000102  
 20 MAMMA1000118 MAMMA1000141 MAMMA1000449  
 MAMMA1000457 MAMMA1000591 MAMMA1000652  
 MAMMA1000681 MAMMA1000986 MAMMA1000994  
 MAMMA1001043 MAMMA1001141 MAMMA1001284  
 MAMMA1001310 MAMMA1001344 MAMMA1001893  
 25 MAMMA1001901 MAMMA1001957 MAMMA1002070  
 MAMMA1002087 MAMMA1002165 MAMMA1002205  
 MAMMA1002224 MAMMA1002633 NT2RM2000241  
 NT2RM2000306 NT2RM2000410 NT2RM2000514  
 NT2RM2001643 NT2RM2001941 NT2RM4000115  
 30 NT2RM4000997 NT2RM4001321 NT2RM4001325  
 NT2RM4001768 NT2RP1000050 NT2RP1000448  
 NT2RP1000903 NT2RP1001563 NT2RP2000479  
 NT2RP2001495 NT2RP2001915 NT2RP2001948  
 NT2RP2002015 NT2RP2002063 NT2RP2002304  
 35 NT2RP2002674 NT2RP2002721 NT2RP2003383  
 NT2RP2003469 NT2RP2003593 NT2RP2003599  
 NT2RP2003655 NT2RP2003664 NT2RP2004179  
 NT2RP2004447 NT2RP2004495 NT2RP2004524  
 NT2RP2004556 NT2RP2004837 NT2RP2005027  
 40 NT2RP2005463 NT2RP2005514 NT2RP2005887  
 NT2RP2006042 NT2RP2006269 NT2RP3000169  
 NT2RP3000460 NT2RP3000481 NT2RP3000645  
 NT2RP3000789 NT2RP3000818 NT2RP3001012  
 NT2RP3001044 NT2RP3001195 NT2RP3001560  
 45 NT2RP3001685 NT2RP3001858 NT2RP3002160  
 NT2RP3002281 NT2RP3002721 NT2RP3002836  
 NT2RP3002958 NT2RP3003076 NT2RP3003354  
 NT2RP3003469 NT2RP3003535 NT2RP3003559  
 NT2RP3003963 NT2RP3004000 NT2RP3004083  
 50 NT2RP3004133 NT2RP3004309 NT2RP3004321  
 NT2RP3004355 NT2RP3004374 NT2RP4001001  
 NT2RP4001879 NT2RP4002451 NT2RP4002715  
 OVARC1000208 OVARC1000298 OVARC1000439  
 OVARC1000775 OVARC1000811 OVARC1000853  
 55 OVARC1001222 OVARC1001727 OVARC1001807  
 OVARC1001833 PLACE1000231 PLACE1000560  
 PLACE1000740 PLACE1000912 PLACE1000914  
 PLACE1000927 PLACE1000986 PLACE1001100

PLACE1001183 PLACE1001229 PLACE1001407  
 PLACE1001536 PLACE1001788 PLACE1002080  
 PLACE1002095 PLACE1002374 PLACE1002518  
 PLACE1003407 PLACE1003428 PLACE1003460  
 5 PLACE1003839 PLACE1003845 PLACE1004028  
 PLACE1004199 PLACE1004282 PLACE1004305  
 PLACE1004482 PLACE1004637 PLACE1005005  
 PLACE1005250 PLACE1005383 PLACE1005410  
 PLACE1005544 PLACE1005569 PLACE1005601  
 10 PLACE1005660 PLACE1005669 PLACE1005725  
 PLACE1005768 PLACE1005927 PLACE1006079  
 PLACE1006093 PLACE1006219 PLACE1006277  
 PLACE1006443 PLACE1006786 PLACE1006809  
 PLACE1007040 PLACE1007096 PLACE1007296  
 15 PLACE1007626 PLACE1007971 PLACE1008469  
 PLACE1008984 PLACE1008985 PLACE1009067  
 PLACE1009196 PLACE1009527 PLACE1009982  
 PLACE1010251 PLACE1011236 PLACE2000219  
 PLACE4000455 SKNMC1000004 SKNMC1000014  
 20 THYRO1000036 THYRO1000099 THYRO1000196  
 THYRO1000795 THYRO1000999 THYRO1001237  
 THYRO1001327 THYRO1001478 THYRO1001495  
 THYRO1001523 THYRO1001702 THYRO1001725  
 Y79AA1000226 Y79AA1000270 Y79AA1000426  
 25 Y79AA1000521 Y79AA1000776 Y79AA1000959  
 Y79AA1001013 Y79AA1001056 Y79AA1001264  
 Y79AA1001328 Y79AA1001427 Y79AA1001430  
 Y79AA1001530 Y79AA1001592 Y79AA1001793  
 Y79AA1001795 Y79M1001803 Y79AA1001863  
 30 Y79AA1002022 Y79AA1002373

[0056] In the example mentioned below, the 254 clones as described above were categorized into three groups according to their maximal value in the ATGpr and the result in the PSORT, which are shown in Table 7-10, 11, 12 (246 clones), and Table 13, 14, 15 (8 clones). In the tables, the name of clone, indicate the name of the clone that was selected by the ATGpr and the PSORT; the name of sequence indicates the name of the 5'-end sequence of the clone on the left; the maximal ATGpr score indicates the maximal ATGpr1 score of the 5'-end sequence shown on the left; and signal indicates the presence of the signal sequence according to the prediction by the PSORT. In addition, the representative sequence is the sequence that has the longest sequence among the cluster in which the 5'-end sequence on the left was included. The maximal ATGpr score and signal on the right indicate the maximal ATGpr1 score of the representative sequence, and the presence of a signal sequence in the representative sequence according to the prediction by the PSORT, respectively. The 170 clones shown in Table 7-10, having the maximal score in the ATGpr1 higher than 0.5, and predicted to possess a signal sequence by the PSORT, are very likely to be full-length and encode a secretory or membrane protein. The 35 clones in Table 11, which have the maximal score in the ATGpr1 0.3 or higher and less than 0.5, and predicted to have a signal sequence, are also as well. And, the 41 clones in Table 12, having the maximal score in the ATGpr1 0 or higher and less than 0.3, and predicted to have a signal sequence, are likely to be full-length and encode a secretory or membrane protein.

[0057] The 8 clones in Table 13 (4 clones), Table 14 (2 clones), and Table 15 (2 clones) have the maximal score in the ATGpr1 0.5 or higher, 0.3 or higher and less than 0.5, and 0 or higher and less than 0.3, respectively, and are predicted to have no signal sequence by the PSORT. However, these clones contain a region that is recognized by the PSORT to be a signal sequence within the representative sequence composing the same cluster. Thus, the clones were judged as a full-length clone which encodes a membrane protein, especially.

[0058] The clones selected by the score in the ATGpr and by the keywords in the top hit data in the SwissProt are likely to encode a secretory or membrane protein, or proteins with functions associated to signal transduction, glyco-protein, transcription, and diseases according to the respective keywords. These 659 clones are shown below. Here, top hit data is defined to be data of known amino acid sequence which is identified to be the most homologous sequence in homology search using the SwissProt.

BNGH41000020 BNGH41000087 BNGH41000091

HEMBA1000006 HEMBA1000121 HEMBN1000128  
 HEMBA1000275 HEMBA1000349 HEMBA1000443  
 HEMBA1000462 HEMBA1000477 HEMBA1000590  
 HEMBA1000634 HEMBA1000671 HEMBA1000732  
 5 HEMBA1000745 HEMBA1000835 HEMBA1000875  
 HEMBA1000907 HEMBA1000940 HEMBA1001184  
 HEMBA1001221 HEMBA1001228 HEMBA1001296  
 HEMBA1001390 HEMBA1001563 HEMBA1001621  
 HEMBA1001878 HEMBA1001886 HEMBA1002048  
 10 HEMBA1002131 HEMBA1002163 HEMBA1002164  
 HEMBA1002167 HEMBA1002178 HEMBA1002195  
 HEMBA1002227 HEMBA1002316 HEMBA1002421  
 HEMBA1002524 HEMBA1002551 HEMBA1002767  
 HEMBA1002985 HEMBA1002992 HEMBA1003047  
 15 HEMBA1003072 HEMBA1003101 HEMBA1003120  
 HEMBA1003230 HEMBA1003315 HEMBA1003392  
 HEMBA1003487 HEMBA1003497 HEMBA1003530  
 HEMBA1003945 HEMBA1004007 HEMBA1004067  
 HEMBA1004085 HEMBA1004250 HEMBA1004391  
 20 HEMBA1004444 HEMBA1004454 HEMBA1004505  
 HEMBA1004785 HEMBA1004797 HEMBA1004952  
 HEMBA1004971 HEMBA1004982 HEMBA1005070  
 HEMBA1005084 HEMBA1005145 HEMBA1005230  
 HEMBA1005246 HEMBA1005267 HEMBA1005337  
 25 HEMBA1005449 HEMBA1005489 HEMBA1005522  
 HEMBA1005545 HEMBA1005698 HEMBA1005913  
 HEMBA1005929 HEMBA1005945 HEMBA1006276  
 HEMBA1006299 HEMBA1006335 HEMBA1006430  
 HEMBA1006482 HEMBA1006517 HEMBA1006544  
 30 HEMBA1006572 HEMBA1006707 HEMBA1006724  
 HEMBA1006749 HEMBA1006770 HEMBA1006902  
 HEMBA1006912 HEMBA1006916 HEMBA1007013  
 HEMBA1007057 HEMBA1007063 HEMBA1007226  
 HEMBA1007241 HEMBA1007291 HEMBA1007332  
 35 HEMBB1000106 HEMBB1000309 HEMBB1000407  
 HEMBB1000447 HEMBB1000542 HEMBB1000567  
 HEMBB1000668 HEMBB1000679 HEMBB1000881  
 HEMBB1001026 HEMBB1001048 HEMBB1001200  
 HEMBB1001573 HEMBB1001847 HEMBB1001959  
 40 HEMBB1002039 HEMBB1002041 HEMBB1002051  
 HEMBB1002120 HEMBB1002302 HEMBB1002427  
 HEMBB1002661 MAMMA1000106 MAMMA1000204  
 MAMMA1000226 MAMMA1000403 MAMMA1000473  
 MAMMA1000496 MAMMA1000528 MAMMA1000591  
 45 MAMMA1000614 MAMMA1000681 MAMMA1000706  
 MAMMA1000788 MAMMA1000810 MAMMA1000814  
 MAMMA1000881 MAMMA1001043 MAMMA1001066  
 MAMMA1001094 MAMMA1001150 MAMMA1001237  
 MAMMA1001418 MAMMA1001532 MAMMA1001609  
 50 MAMMA1001615 MAMMA1001623 MAMMA1001634  
 MAMMA1001893 MAMMA1001957 MAMMA1001978  
 MAMMA1002070 MAMMA1002080 MAMMA1002091  
 MAMMA1002095 MAMMA1002128 MAMMA1002142  
 MAMMA1002165 MAMMA1002234 MAMMA1002586  
 55 MAMMA1002633 MAMMA1003126 NT2RM1000407  
 NT2RM1000462 NT2RM1000542 NT2RM1000580  
 NT2RM1000789 NT2RM1000855 NT2RM1000858  
 NT2RM1000899 NT2RM2000410 NT2RM2000423

NT2RM2000497 NT2RM2000565 NT2RM2000582  
 NT2RM2000589 NT2RM2000622 NT2RM2000632  
 NT2RM2000773 NT2RM2001126 NT2RM2001558  
 NT2RM2001626 NT2RM2001738 NT2RM2001767  
 5 NT2RM2001792 NT2RM2001818 NT2RM2001902  
 NT2RM2001939 NT2RM2001941 NT2RM4000100  
 NT2RM4000198 NT2RM4000284 NT2RM4000295  
 NT2RM4000326 NT2RM4000417 NT2RM4000444  
 NT2RM4000587 NT2RM4000593 NT2RM4000648  
 10 NT2RM4000761 NT2RM4000965 NT2RM4001377  
 NT2RM4001735 NT2RM4001843 NT2RM4002352  
 NT2RP1000002 NT2RP1000050 NT2RP1000181  
 NT2RP1000239 NT2RP1000261 NT2RP1000271  
 NT2RP1000300 NT2RP1000325 NT2RP1000465  
 15 NT2RP1000468 NT2RP1000551 NT2RP1000579  
 NT2RP1000613 NT2RP1000679 NT2RP1000740  
 NT2RP1000981 NT2RP1001004 NT2RP1001020  
 NT2RP1001031 NT2RP2000092 NT2RP2000178  
 NT2RP2000240 NT2RP2000394 NT2RP2000447  
 20 NT2RP2000514 NT2RP2000533 NT2RP2000610  
 NT2RP2000616 NT2RP2000649 NT2RP2000663  
 NT2RP2000694 NT2RP2000712 NT2RP2000739  
 NT2RP2000818 NT2RP2000903 NT2RP2001200  
 NT2RP2001223 NT2RP2001276 NT2RP2001388  
 25 NT2RP2001469 NT2RP2001480 NT2RP2001495  
 NT2RP2001514 NT2RP2001529 NT2RP2001538  
 NT2RP2001562 NT2RP2001662 NT2RP2001755  
 NT2RP2001769 NT2RP2001817 NT2RP2001878  
 NT2RP2001903 NT2RP2001921 NT2RP2001948  
 30 NT2RP2001956 NT2RP2002063 NT2RP2002188  
 NT2RP2002232 NT2RP2002304 NT2RP2002409  
 NT2RP2002510 NT2RP2002527 NT2RP2002533  
 NT2RP2002564 NT2RP2002824 NT2RP2002942  
 NT2RP2002974 NT2RP2002976 NT2RP2003042  
 35 NT2RP2003138 NT2RP2003179 NT2RP2003210  
 NT2RP2003302 NT2RP2003369 NT2RP2003390  
 NT2RP2003469 NT2RP2003545 NT2RP2003593  
 NT2RP2003655 NT2RP2003664 NT2RP2003931  
 NT2RP2003940 NT2RP2003950 NT2RP2004069  
 40 NT2RP2004108 NT2RP2004141 NT2RP2004205  
 NT2RP2004447 NT2RP2004606 NT2RP2004648  
 NT2RP2004670 NT2RP2004794 NT2RP2004847  
 NT2RP2005069 NT2RP2005163 NT2RP2005181  
 NT2RP2005247 NT2RP2005378 NT2RP2005391  
 45 NT2RP2005425 NT2RP2005535 NT2RP2005541  
 NT2RP2005597 NT2RP2005632 NT2RP2005666  
 NT2RP2005774 NT2RP2005878 NT2RP2005883  
 NT2RP2005941 NT2RP2005994 NT2RP2006004  
 NT2RP2006042 NT2RP2006092 NT2RP2006099  
 50 NT2RP2006134 NT2RP2006269 NT2RP2006512  
 NT2RP3000011 NT2RP3000022 NT2RP3000059  
 NT2RP3000063 NT2RP3000125 NT2RP3000148  
 NT2RP3000171 NT2RP3000172 NT2RP3000201  
 NT2RP3000232 NT2RP3000304 NT2RP3000378  
 55 NT2RP3000427 NT2RP3000436 NT2RP3000444  
 NT2RP3000481 NT2RP3000616 NT2RP3000645  
 NT2RP3000652 NT2RP3000676 NT2RP3000677  
 NT2RP3000721 NT2RP3000820 NT2RP3000838

NT2RP3000871 NT2RP3000907 NT2RP3000921  
 NT2RP3001012 NT2RP3001061 NT2RP3001159  
 NT2RP3001170 NT2RP3001195 NT2RP3001240  
 NT2RP3001271 NT2RP3001322 NT2RP3001388  
 5 NT2RP3001542 NT2RP3001560 NT2RP3001592  
 NT2RP3001650 NT2RP3001738 NT2RP3001754  
 NT2RP3001976 NT2RP3002015 NT2RP3002160  
 NT2RP3002286 NT2RP3002311 NT2RP3002324  
 NT2RP3002342 NT2RP3002353 NT2RP3002409  
 10 NT2RP3002411 NT2RP3002448 NT2RP3002571  
 NT2RP3002664 NT2RP3002737 NT2RP3002738  
 NT2RP3002790 NT2RP3002836 NT2RP3002887  
 NT2RP3002900 NT2RP3002958 NT2RP3002983  
 NT2RP3003000 NT2RP3003076 NT2RP3003354  
 15 NT2RP3003448 NT2RP3003473 NT2RP3003527  
 NT2RP3003532 NT2RP3003614 NT2RP3003729  
 NT2RP3003849 NT2RP3003874 NT2RP3003939  
 NT2RP3004025 NT2RP3004067 NT2RP3004075  
 NT2RP3004090 NT2RP3004119 NT2RP3004130  
 20 NT2RP3004133 NT2RP3004202 NT2RP3004294  
 NT2RP3004309 NT2RP3004345 NT2RP3004406  
 NT2RP3004481 NT2RP3004552 NT2RP3004557  
 NT2RP3004625 NT2RP3004640 NT2RP3004647  
 NT2RP4000108 NT2RP4000634 NT2RP4000962  
 25 NT2RP4001009 NT2RP4001467 NT2RP4001877  
 NT2RP4001879 NT2RP4002187 NT2RP4002451  
 NT2RP4002750 OVARC1000003 OVARC1000090  
 OVARC1000105 OVARC1000137 OVARC1000255  
 OVARC1000275 OVARC1000307 OVARC1000313  
 30 OVARC1000331 OVARC1000410 OVARC1000439  
 OVARC1000467 OVARC1000529 OVARC1000553  
 OVARC1000873 OVARC1000916 OVARC1000956  
 OVARC1000995 OVARC1001030 OVARC1001049  
 OVARC1001086 OVARC1001132 OVARC1001163  
 35 OVARC1001222 OVARC1001260 OVARC1001336  
 OVARC1001338 OVARC1001569 OVARC1001570  
 OVARC1001596 OVARC1001607 OVARC1001725  
 OVARC1001952 OVARC1001991 OVARC1002058  
 OVARC1002178 PLACE1000033 PLACE1000258  
 40 PLACE1000442 PLACE1000740 PLACE1000907  
 PLACE1001016 PLACE1001114 PLACE1001123  
 PLACE1001231 PLACE1001340 PLACE1001401  
 PLACE1001407 PLACE1001464 PLACE1001500  
 PLACE1001516 PLACE1001564 PLACE1001655  
 45 PLACE1001795 PLACE1001836 PLACE1001918  
 PLACE1001949 PLACE1002080 PLACE1002095  
 PLACE1002153 PLACE1002329 PLACE1002355  
 PLACE1002374 PLACE1002547 PLACE1002726  
 PLACE1002905 PLACE1002911 PLACE1002967  
 50 PLACE1003135 PLACE1003163 PLACE1003428  
 PLACE1003438 PLACE1003460 PLACE1003529  
 PLACE1003573 PLACE1003598 PLACE1003644  
 PLACE1003737 PLACE1003772 PLACE1003852  
 PLACE1004078 PLACE1004166 PLACE1004168  
 55 PLACE1004279 PLACE1004441 PLACE1004450  
 PLACE1004482 PLACE1004492 PLACE1004519  
 PLACE1004520 PLACE1004630 PLACE1004648  
 PLACE1004816 PLACE1004887 PLACE1005003



PLACE1005031 PLACE1005239 PLACE1005383  
 PLACE1005426 PLACE1005519 PLACE1005539  
 PLACE1005544 PLACE1005569 PLACE1005682  
 PLACE1005736 PLACE1005745 PLACE1005815  
 5 PLACE1005878 PLACE1005927 PLACE1006071  
 PLACE1006073 PLACE1006208 PLACE1006277  
 PLACE1006290 PLACE1006443 PLACE1006515  
 PLACE1006716 PLACE1006959 PLACE1007028  
 PLACE1007077 PLACE1007081 PLACE1007096  
 10 PLACE1007296 PLACE1007591 PLACE1007702  
 PLACE1007845 PLACE1007881 PLACE1008282  
 PLACE1008297 PLACE1008359 PLACE1008469  
 PLACE1008549 PLACE1008657 PLACE1008716  
 PLACE1008744 PLACE1008984 PLACE1008985  
 15 PLACE1009279 PLACE1009527 PLACE1009546  
 PLACE1009600 PLACE1009735 PLACE1010011  
 PLACE1010078 PLACE1010081 PLACE1010251  
 PLACE1010445 PLACE1010713 PLACE1010784  
 PLACE1010827 PLACE1010968 PLACE1011045  
 20 PLACE1011116 PLACE1011181 PLACE1011236  
 PLACE1011364 PLACE1011407 PLACE1011516  
 PLACE1011708 PLACE1011824 PLACE1011978  
 PLACE2000118 PLACE3000181 PLACE3000213  
 PLACE4000354 SKNMC1000014 SKNMC1000082  
 25 THYRO1000061 THYRO1000196 THYRO1000400  
 THYRO1000580 THYRO1000584 THYRO1000678  
 THYRO1000776 THYRO1000795 THYRO1000846  
 THYRO1000866 THYRO1000956 THYRO1000964  
 THYRO1001063 THYRO1001071 THYRO1001102  
 30 THYRO1001113 THYRO1001128 THYRO1001205  
 THYRO1001242 THYRO1001266 THYRO1001456  
 THYRO1001457 THYRO1001471 THYRO1001478  
 THYRO1001529 THYRO1001593 THYRO1001608  
 THYRO1001641 THYRO1001700 THYRO1001702  
 35 THYRO1001770 THYRO1001803 Y79AA1000030  
 Y79AA1000127 Y79AA1000207 Y79AA1000270  
 Y79AA1000426 Y79AA1000750 Y79AA1000777  
 Y79AA1000876 Y79AA1000888 Y79AA1000967  
 Y79AA1001062 Y79AA1001090 Y79AA1001212  
 40 Y79AA1001272 Y79AA1001426 Y79AA1001523  
 Y79AA1001727 Y79AA1001787 Y79AA1001799  
 Y79AA1001803 Y79AA1001863 Y79AA1002058  
 Y79AA1002121 Y79AA1002129 Y79AA1002213  
 Y79AA1002334 Y79AA1002376 Y79AA1002378  
 45 Y79AA1002381 NT2RP2006580

**[0059]** Among the clones, the following 83 clones are identical to the clones selected by the score in the ATGpr and the prediction by the PSORT for the existence of a signal sequence.

50 HEMBA1000907 NT2RM2000410 PLACE1000740  
 HEMBA1002164 NT2RM2001941 PLACE1001407  
 HEMBA1002421 NT2RP1000050 PLACE1002080  
 HEMBA1003101 NT2RP2001495 PLACE1002095  
 HEMBA1004797 NT2RP2001948 PLACE1002374  
 55 HEMBA1006335 NT2RP2002063 PLACE1003428  
 HEMBA1006572 NT2RP2002304 PLACE1003460  
 HEMBA1006707 NT2RP2003469 PLACE1004482  
 HEMBA1006902 NT2RP2003593 PLACE1005383

HEMBA1007013 NT2RP2003655 PLACE1005544  
 HEMBB1000447 NT2RP2003664 PLACE1005569  
 HEMBB1000567 NT2RP2004447 PLACE1005927  
 HEMBB1001200 NT2RP2006042 PLACE1006277  
 5 HEMBB1002427 NT2RP2006269 PLACE1006443  
 MAMMA1000591 NT2RP3000481 PLACE1007096  
 MAMMA1000681 NT2RP3000645 PLACE1007296  
 MAMMA1001043 NT2RP3001012 PLACE1008469  
 MAMMA1001893 NT2RP3001195 PLACE1008984  
 10 MAMMA1001957 NT2RP3001560 PLACE1008985  
 MAMMA1002070 NT2RP3002160 PLACE1009527  
 MAMMA1002165 NT2RP3002836 PLACE1010251  
 MAMMA1002633 NT2RP3002958 PLACE1011236  
 NT2RP3003076 SKNMC1000014  
 15 NT2RP3003354 THYPO1000196  
 NT2RP3004133 THYRO1000795  
 NT2RP3004309 THYRO1001478  
 NT2RP4001879 THYRO1001702  
 NT2RP4002451 Y79AA1000270  
 20 OVARC1000439 Y79AA1000426  
 OVARC1001222 Y79AA1001803  
 Y79AA1001863

25 **[0060]** The 446 clones in Table 16, 17, 18, 19, and 20, and NT2RP2006580 are predicted to encode a secretory or membrane protein. Among them, 77 clones were identical to the clones selected by the score in the ATGpr and the prediction by the PSORT for the existence of a signal sequence (overlapping with any of the 254 clones listed in Table 7-15). Besides, many clones were turned out to be identical to the clones selected as a protein associated with a glycoprotein. Also, there were clones identical to those selected as a protein associated with a disease.

30 **[0061]** The 243 clones in Table 21 are predicted to encode a glycoprotein. Among them, 53 clones were identical to those selected by the score in the ATGpr and the prediction by the PSORT for the existence of a signal sequence. And, many clones were turned out to be identical to the clones selected as a secretory or membrane protein. Moreover, there were clones identical to those selected as a protein associated with a disease.

**[0062]** The 51 clones in Table 22 are predicted to encode a protein associated to signal transduction.

**[0063]** The 130 clones in Table 23 are predicted to encode a protein associated to transcription.

35 **[0064]** The 17 clones in Table 24 are predicted to encode a protein associated with diseases.

**[0065]** In these clones, 532 clones have the maximal ATGpr1 score of 0.5 or higher (Table 25). 60 clones have the maximal ATGpr1 score of 0.3 or higher and less than 0.5 (Table 26 and NT2RP2006580). And 67 clones were with the maximal ATGpr1 score of 0 or higher and less than 0.3 (Table 27).

40 **[0066]** 532 clones shown in Table 25, each having the maximal score in the ATGpr1 0.5 or higher, are very likely to be full-length and encode a secretory or membrane protein, or proteins associated to signal transduction, glycoprotein, transcription, or diseases. 59 clones in Table 26 and NT2RP2006580, which have the maximal score in the ATGpr1 0.3 or higher and less than 0.5, are likely to be full-length and encode a secretory or membrane protein, or proteins associated to signal transduction, glycoprotein, transcription, or diseases. 67 clones in Table 27, having the maximal score in the ATGpr1 0 or higher and less than 0.3, are still likely to be full-length and encode a secretory or membrane protein, or proteins associated to signal transduction, glycoprotein, transcription, or diseases.

45 **[0067]** This is the method for selecting the cDNA clones predicted to encode secretory and/or transmembrane proteins, glycoprotein-associated proteins, signal transduction-associated proteins, transcription-associated proteins, or disease-associated proteins on the basis of the partial sequences (5' sequences).

50 **[0068]** The polynucleotide of the present invention encodes an amino acid sequence of a functional protein such as a secretory or membrane protein, or a protein associated to signal transduction, glycoprotein, transcription, or diseases. Since the protein has the complete amino acid sequence, it is possible to analyze its biological activity by expressing the protein as a recombinant protein using an appropriate expression system, or by raising and using an antibody which specifically recognizes it.

55 **[0069]** It is possible to analyze the biological activity of a secretory protein or a membrane protein, or proteins associated to signal transduction, glycoprotein, or transcription, based on the methods in "Gene Transcription" (Hames B. D., and Higgins S.J. edit, (1993)), "Glycobiology" (Fukuda M., and Kobata A. edit, (1993)), "Growth Factors" (McKay I., and Leigh I. edit, (1993)), "Extracellular Matrix" (Haralson M.A., and Hassell J.R. edit, (1995)), "Transcription Factors" (Latchman D.S. edit, (1993)), "Signal Transduction" (Milligans G. edit, (1992)), featured in "The Practical Approach

Series" (IRL PRESS), or "Signal Transduction Protocols" (Kendall D.A., and Hill S.J. edit, (1995), "Glycoprotein Analysis in Biomedicine" (Hounsell E.F. edit, (1993)), featured in "Method in Molecular Biology" (Humana Press).

[0070] As to a protein associated with a disease, it is possible to perform a functional analysis as described above, but also possible to analyze correlation between the expression or the activity of the protein and a certain disease by using a specific antibody that recognizes the protein. Alternatively, it is possible to utilize the database Online Mendelian Inheritance in Man (OMIM) (<http://www.ncbi.nlm.nih.gov/Omim/>), which is a database of human genes and diseases, to analyze the protein. New information is constantly being deposited in the OMIM database. Therefore, it is possible for one skilled in the art to find a new relationship between a particular disease and a gene of the present invention in the updated database.

[0071] Proteins associated with diseases are useful in drug development as they can be utilized as a diagnostic marker, a drug that regulates the level of their expression and activities, or a target of gene therapy. Also, as for a secretory protein, membrane protein, or proteins associated with signal transduction, glycoprotein, or transcription, search of the OMIM with the keywords mentioned below revealed that the proteins are associated with many diseases. Also, relationship between a proteins associated to signal transduction or transcription and diseases is reported in "Transcription Factor Research-1999" (Fujii, Tamura, Morohashi, Kageyama, and Satake edit, (1999) Jikken-Igaku Zoukan, Vol.17, No.3), and "Gene Medicine" ((1999) Vol.3, No.2). Thus, not only a protein associated with diseases, but also a secretory protein, membrane protein, or protein associated with signal transduction, glycoprotein, or transcription is involved in diseases, suggesting these proteins also are very important as a target in medical industry.

[0072] Keywords used in the search of the OMIM

- (1) secretion protein
- (2) membrane protein
- (3) channel
- (4) extracellular matrix
- (5) receptor
- (6) glycoprotein
- (7) protein kinase
- (8) calmodulin kinase
- (9) transcription factor

[0073] Shown in the search result are only the accession numbers in the OMIM. Using the number, data showing the relationship between a disease and a gene or protein can be seen. The OMIM data has been renewed everyday.

#### 1) Secretion protein

268 entries found, searching for "secretion protein"

104760, 176860, 160900, 107400, 118910, 139320, 603850, 147572, 176880, 600946, 603215, 157147, 600174, 151675, 170280, 179512, 179513, 138120, 179509, 246700, 179510, 600626, 179511, 600998, 109270, 601489, 154545, 179490, 185860, 603216, 122559, 601746, 147290, 602672, 146770, 603062, 179508, 131230, 601591, 602421, 139250, 167805, 167770, 600041, 600564, 118825, 601146, 300090, 600753, 601652, 600759, 600768, 602434, 182590, 603166, 308230, 602534, 603489, 107470, 150390, 104610, 173120, 158106, 143890, 306900, 308700, 134797, 137350, 227500, 176300, 107730, 600760, 138079, 120180, 120160, 120150, 124092, 138160, 101000, 227600, 600509, 601199, 142410, 104311, 193400, 201910, 107300, 122560, 272800, 217000, 590050, 147670, 133170, 176730, 300300, 134370, 274600, 120140, 162151, 158070, 152790, 120120, 106100, 300200, 192340, 190160, 138040, 147470, 147620, 173350, 147380, 152200, 152760, 157145, 153450, 264080, 113811, 600937, 600840, 188545, 202110, 600514, 186590, 603372, 136435, 137241, 252800, 214500, 207750, 138850, 139191, 142640, 138130, 189907, 603692, 600633, 603355, 107270, 600377, 147892, 232200, 600281, 232800, 602358, 137035, 601771, 601769, 253200, 601933, 118444, 600270, 120700, 600945, 603732, 147660, 600761, 172400, 600823, 600877, 130080, 171060, 107740, 307800, 602843, 130660, 152780, 124020, 601124, 601340, 601604, 601610, 171050, 312060, 232700, 300159, 142703, 600734, 125255, 168450, 123812, 188540, 147940, 188450, 600839, 182452, 188400, 182280, 176760, 263200, 600264, 188826, 252650, 601185, 162641, 137216, 601398, 601538, 118888, 118445, 601745, 190180, 601922, 182098, 602008, 147440, 602384, 600031, 109160, 602663, 151670, 602682, 602730, 602779, 146880, 603061, 142704, 603140, 106150, 600732, 153620, 603318, 139392, 600042, 102200, 603493, 182100, 264300, 603795, 184600

#### 2) Membrane protein

1017 entries found, searching for "membrane protein"

130500, 305360, 153330, 173610, 170995, 109270, 170993, 309060, 120920, 602333, 133740, 133710, 602690,

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### 3) Channel

272 entries found, searching for "channel"

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### 4) Extracellular matrix

167 entries found, searching for "extracellular matrix"

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### 5) Receptor (including membrane proteins, and also including transcription factors, since nuclear proteins were not excluded in the search)

1606 entries found, searching for "receptor"

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## 6) Glycoprotein

438 entries found, searching for "glycoprotein"

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#### 7) Protein kinase (a member of signal transduction)

729 entries found, searching for "protein kinase"

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## 8) Calmodulin kinase (a member of signal transduction)

35 entries found, searching for "calmodulin binding"

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## 9) Transcription factor

717 entries found, searching for "transcription factor"

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187011, 102600, 180380, 162080, 603450, 142967, 602301, 126375, 603372, 603355, 164720, 603250, 167409,  
167415, 602897, 601565, 185250, 182138, 601851, 600749, 601575, 194548,  
154500, 601365, 194541, 601621, 601623, 601531, 600790, 194355, 123830, 123812, 154540, 601415, 143055,  
601386, 194550, 186930, 131290, 601320, 601620, 601754, 601313, 184430, 182900, 182500, 600725, 147870,  
20 154365, 116953, 601297, 601296, 601265, 600796, 120436, 601644, 601930, 601643, 230200, 601645, 601972,  
600861, 602009, 601172, 601158, 601646, 180630, 600821, 118440, 601656, 601647, 150200, 601125, 601671,  
141850, 116899, 600697, 109270, 202110, 150570, 601108, 191339, 601063, 109691, 180240, 203100, 151430,  
179710, 111000, 176797, 238600, 104311, 240300, 125255, 600423, 158070, 602439, 600324, 112261, 243305,  
602474, 174762, 600613, 602539, 138890, 138720, 114550, 173865, 602582, 602584, 173510, 600250, 602627,  
25 173325, 602635, 246530, 172425, 600193, 602691, 600188, 170998, 152790,  
168468, 256540, 225250, 600848, 143400, 168461, 262600, 168360, 601912, 602951, 600017, 230000, 266600,  
602981, 272800, 109150, 102200, 603025, 603026, 603109, 167050, 603127, 603128, 165240, 230400, 313700,  
164975, 164875, 602017, 115500, 235800, 164873, 602110, 164785, 164772, 312865, 603296, 600542, 164740,  
602125, 309801, 602148, 300007, 306955, 603368, 116940, 602181, 603416, 126650, 163920, 300024, 603437,  
30 602209, 603576, 603607, 305435, 600944, 180410, 303630, 159557, 301870, 132810, 100790, 603849, 603862,  
603881,

[0074] There are several methods for analyzing the expression levels of genes associated with diseases. Differences  
in gene expression levels between diseased and normal tissues are studied by the analytical methods, for example,  
35 Northern hybridization and differential display. Other examples include a method with high-density cDNA filter, a method  
with DNA microarray and methods with PCR amplification (Experimental Medicine, Vol.17, No. 8, 980-1056 (1999);  
Cell Engineering (additional volume) DNA Microarray and Advanced PCR Methods, Muramatsu & Naba (eds.), Shu-  
junsya). The levels of gene expression between diseased tissues and normal tissues can be studied by any of these  
analytical methods. When explicit difference in expression level is observed for a gene, it can be concluded that the  
40 gene is closely associated with a disease or disorder. Instead of diseased tissues, cultured cells can be used for the  
assessment. Similarly, when gene expression is explicitly different between normal cells and cells reproducing disease-  
associated specific features, it can be concluded that the gene is closely associated with a disease or disorder. When  
the expression levels of genes are evidently varied during major cellular events (such as differentiation and apoptosis),  
the genes are involved in the cellular events and accordingly are candidates for disease- and/or disorder-associated  
45 genes. Further, genes exhibiting tissue-specific expression are genes playing important parts in the tissue functions  
and, therefore, can be candidates for genes associated with diseases and/or disorders affecting the tissues.

[0075] For example, non-enzymic protein glycation reaction is believed to be a cause for a variety of chronic diabetic  
complications. Accordingly, in endothelial cells, genes, of which expression levels are elevated or decreased in a gly-  
cated protein-dependent manner, are associated with diabetic complications caused by glycated proteins (Diabetes  
50 1996, 45 (Suppl. 3), S67-S72; Diabetes 1997, 46 (Suppl. 2), S19-S25). The onset of rheumatoid arthritis is thought to  
be involved in the proliferation of synovial cells covering inner surfaces of joint cavity and in inflammatory reaction  
resulted from the action of cytokines produced by leukocytes infiltrating into the joint synovial tissues (Rheumatism  
Information Center, <http://www.rheuma-net.or.jp/>). Recent studies have also revealed that tissue necrosis factor (TNF)-  
55  $\alpha$  participates in the onset (Current opinion in immunology 1999, 11, 657-662). When the expression of a gene exhibits  
responsiveness to the action of TNF on synovial cells, the gene is considered to be involved in rheumatoid arthritis.  
Many genes acting at the downstream of TNF- and IL-1, among inflammation-associated cytokines have been previ-  
ously identified. The respective stimulations are transduced through independent pathways of signaling cascade. There  
exists another signaling cascade for both stimulations, wherein NF- $\kappa$ B is a common transducing molecule shared by

the two stimulations (J. Leukoc. Biol., 1994, 56(5): 542-547). It has also been revealed that many inflammation-associated genes, including IL-2, IL-6 and G-CSF, are varied in the expression levels in response to the signal through the common pathway (Trends Genet. 1999, 15(6): 229-235). It is assumed that genes of which expression levels are varied in response to the stimulation of TNF- or IL-1, also participate in inflammation. Genes associated with neural differentiation can be candidates for causative genes for neurological diseases as well as candidates for genes usable for treating the diseases.

**[0076]** Clones exhibiting differences in the expression levels thereof can be selected by using gene expression analysis. The selection comprises, for example; analyzing cDNA clones by using high-density cDNA filter; and statistically treating the multiple signal values (signal values of radioisotope in the radiolabeled probes or values obtained by measuring fluorescence intensities emitted from the fluorescent labels) for the respective clones by two-sample t-test, where the signal values are determined by multiple experiments of hybridization. The clones of interest are selectable based on the statistically significant differences in the signal distribution at  $p < 0.05$ . However, selectable clones with significant difference in the expression levels thereof may be changed depending on the partial modification of statistical treatment. For example, the clones may be selected by conducting statistical treatment with two-sample t-test at  $p < 0.01$ ; or genes exhibiting more explicit differences in the expression levels thereof can be selected by performing statistical treatment with a pre-determined cut-off value for the significant signal difference. An alternative method is that the expression levels are simply compared with each other, and then, the clones of interest are selected based on the ratio of the expression levels thereof.

**[0077]** Clones exhibiting differences in the expression levels can also be selected by comparing the expression levels by PCR analysis, for example, by using the method of determining the band intensities representing the amounts of PCR products with ethidium bromide staining; the method of determining the radioisotopic signal values or fluorescence intensities of the PCR products when radio-labeled or fluorescence-labeled primers; or the method of determining the values of radioisotope signals or fluorescence intensities of the probes hybridized to the PCR products when radio-labeled or fluorescence-labeled probes, respectively, are used in the hybridization. If the expression level ratios obtained in multiple PCR experiments are constantly at least 2-fold, such a clone can be judged to exhibit the difference in the expression level. When the ratios are several-fold or not less than 10-fold, the clone can be selected as a gene exhibiting the explicit difference in the expression level.

**[0078]** A survey of genes of which expression levels are varied specifically to the glycosylated protein in the endothelial cells revealed three genes with elevated expression levels, NT2RP2001538, NT2RP4001001 and Y79AA1000967. These clones are genes associated with diabetes.

**[0079]** A survey of genes of which expression levels are varied in response to TNF. (Tumor Necrosis Factor- $\alpha$ ) in the primary cell culture of synovial tissue detected the following clones with elevated expression levels in the presence of TNF.:

BNGH41000020, HEMBA1000349, HEMBA1000634, HEMBA1000671, HEMBA1000835, HEMBA1000962, HEMBA1002178, HEMBA1002195, HEMBA1002239, HEMBA1002420, HEMBA1002524, HEMBA1002992, HEMBA1003315, HEMBA1003392, HEMBA1003487, HEMBA1003602, HEMBA1004067, HEMBA1004797, HEMBA1005337, HEMBA1005489, HEMBA1006916, HEMBB1000668, HEMBB1000905, HEMBB1001547, HEMBB1001573, HEMBB1002041, HEMBB1002663, MAMMA1000652, MAMMA1000810, MAMMA1001634, MAMMA1002091, MAMMA1002234, NT2RM2000306, NT2RM4000417, NT2RP1000002, NT2RP1000181, NT2RP1000740, NT2RP2000694, NT2RP2001921, NT2RP2002527, NT2RP2004495, NT2RP2004606, NT2RP2005163, NT2RP2005463, NT2RP2006134, NT2RP3000171, NT2RP3000652, NT2RP3001195, NT2RP3001976, NT2RP3003473, NT2RP3003874, NT2RP3004090, NT2RP3004294, NT2RP3004557, NT2RP3004647, NT2RP4000108, NT2RP4001001, NT2RP4001877, OVARC1000090, OVARC1000105, OVARC1000275, OVARC1000439, OVARC1001607, PLACE1000740, PLACE1000927, PLACE1001016, PLACE1001100, PLACE1001464, PLACE1001500, PLACE1001918, PLACE1002095, PLACE1002547, PLACE1003644, PLACE1004519, PLACE1005031, PLACE1005410, PLACE1005736, PLACE1006219, PLACE1006809, PLACE1008716, PLACE1010081, THYRO1001770, Y79AA1000127, Y79M1000207, Y79AA1000270, Y79AA1000876, Y79AA1001013, Y79AA1001264, Y79AA1001272, Y79AA1001328, Y79AA1001430, Y79AA1001530, Y79AA1001799

**[0080]** Clones with decreased expression levels in the presence of TNF are NT2RM4000326, NT2RP1000300, NT2RP2000514, NT2RP2001755, NT2RP2006042, NT2RP3000481, NT2RP3002790. These clones are candidates for rheumatoid arthritis-associated genes.

**[0081]** A survey of genes of which expression levels are varied in response to TNF. (Tumor Necrosis Factor- $\alpha$ ) or IL-1. (Interleukin-1  $\beta$ ) in a human T cell strain, Jurkat cell, revealed the following clones with elevated expression levels in the presence of TNF.:

MAMMA1000141, MAMMA1000788, MAMMA1001237, MAMMA1002070, NT2RM2000582, NT2RM2002109, NT2RP1000679, NT2RP2003664, NT2RP2004108, NT2RP2005597, NT2RP3001592, NT2RP3002738, NT2RP3004133, NT2RP3004294, NT2RP3004321, NT2RP3004557, PLACE1002547, PLACE1003573,

PLACE1004305, PLACE1008744, PLACE1010011, PLACE1010713, PLACE1011181, Y79AA1000776, Y79AA1002129

[0082] The survey also revealed a clone of which expression level was decreased in the presence of TNF. The clone is PLACE1002070. The same survey further revealed the clones of which expression levels were elevated in the presence of IL-1.. The clones are MAMMA1000614, MAMMA1001237, NT2RM2000514 and NT2RP3001159. These clones are genes associated with inflammation.

[0083] A survey of genes of which expression levels are varied in response to the stimulation for inducing cell differentiation (stimulation using retinoic acid (RA) or using RA/inhibitor (inhibitor for cell division)) in cultured cells of neural strain, NT2, revealed the following clones with elevated expression levels in the presence of RA:

10 HEMBA1000121, HEMBA1000275, HEMBA1000300, HEMBA1000634, HEMBA1000671, HEMBA1000875, HEMBA1001184, HEMBA1001390, HEMBA1001886, HEMBA1002163, HEMBA1002227, HEMBA1002420, HEMBA1002421, HEMBA1003072, HEMBA1003120, HEMBA1003294, HEMBA1003497, HEMBA1004007, HEMBA1004110, HEMBA1004391, HEMBA1004444, HEMBA1005230, HEMBA1005246, HEMBA1005267, HEMBA1005489, HEMBA1005913, HEMBA1006299, HEMBA1006357, HEMBA1006517, HEMBA1006544, 15 HEMBA1006658, HEMBA1006749, HEMBA1007063, HEMBA1007241, HEMBB1000447, HEMBB1000542, HEMBB1000567, HEMBB1000642, HEMBB1000668, HEMBB1001026, HEMBB1001847, HEMBB1002051, HEMBB1002120, HEMBB1002228, HEMBB1002693, MAMMA1000106, MAMMA1000141, MAMMA1000473, MAMMA1000528, MAMMA1000810, MAMMA1000881, MAMMA1001634, MAMMA1001957, MAMMA1002205, MAMMA1002224, NT2RM2000423, NT2RM2000497, NT2RM2000582, NT2RM2001126, NT2RM2001902, 20 NT2RM4000198, NT2RM4000284, NT2RM4000593, NT2RM4001321, NT2RP1000002, NT2RP1000050, NT2RP1000181, NT2RP1000261, NT2RP1000465, NT2RP1000468, NT2RP1000579, NT2RP1000679, NT2RP2000092, NT2RP2000479, NT2RP2000610, NT2RP2000663, NT2RP2000694, NT2RP2000903, NT2RP2001388, NT2RP2001538, NT2RP2001878, NT2RP2002015, NT2RP2002304, NT2RP2002721, NT2RP2002824, NT2RP2002942, NT2RP2002974, NT2RP2002976, NT2RP2003179, NT2RP2003302, 25 NT2RP2003383, NT2RP2003469, NT2RP2003664, NT2RP2003940, NT2RP2004069, NT2RP2004108, NT2RP2004524, NT2RP2004556, NT2RP2004670, NT2RP2005069, NT2RP2005247, NT2RP2005425, NT2RP2005463, NT2RP2005514, NT2RP2005535, NT2RP2005541, NT2RP2005774, NT2RP2005878, NT2RP2005883, NT2RP2005887, NT2RP2006099, NT2RP2006134, NT2RP3000011, NT2RP3000125, NT2RP3000171, NT2RP3000232, NT2RP3000460, NT2RP3000481, 30 NT2RP3000652, NT2RP3000677, NT2RP3000818, NT2RP3000820, NT2RP3001044, NT2RP3001061, NT2RP3001170, NT2RP3001240, NT2RP3001322, NT2RP3001388, NT2RP3001542, NT2RP3001592, NT2RP3001976, NT2RP3002790, NT2RP3002900, NT2RP3002983, NT2RP3003000, NT2RP3003354, NT2RP3003532, NT2RP3003729, NT2RP3003874, NT2RP3003939, NT2RP3004025, NT2RP3004083, NT2RP3004090, NT2RP3004130, NT2RP3004202, NT2RP3004294, NT2RP3004640, NT2RP4000108, 35 NT2RP4000634, NT2RP4002451, NT2RP4002715, OVARC1000090, OVARC1000208, OVARC1000275, OVARC1000553, OVARC1000775, OVARC1000853, OVARC1000873, OVARC1000916, OVARC1000995, OVARC1001030, OVARC1001049, OVARC1001132, OVARC1001596, OVARC1002178, PLACE1000258, PLACE1000442, PLACE1000927, PLACE1000986, PLACE1001100, PLACE1001123, PLACE1001795, PLACE1002518, PLACE1002547, PLACE1002967, PLACE1003407, PLACE1003428, PLACE1003644, 40 PLACE1003839, PLACE1004078, PLACE1004441, PLACE1004450, PLACE1005669, PLACE1005682, PLACE1005736, PLACE1005768, PLACE1005815, PLACE1006073, PLACE1006208, PLACE1007296, PLACE1007626, PLACE1008282, PLACE1008984, PLACE1008985, PLACE1010445, PLACE1011708, PLACE1011978, PLACE4000455, SKNMC1000004, THYRO1000036, THYRO1000580, THYRO1000776, THYRO1000999, THYRO1001063, THYRO1001128, THYRO1001205, 45 THYRO1001327, THYRO1001523, THYRO1001725, THYRO1001770, Y79AA1000207, Y79AA1000226, Y79AA1000270, Y79AA1001056, Y79AA1001062, Y79AA1001090, Y79AA1001727, Y79AA1002213, Y79AA1002381

[0084] The survey also revealed the clones of which expression levels were decreased in the presence of RA. The clones are BNGH41000020, HEMBA1005070, NT2RP2005027, NT2RP3003473 and Y79AA1002376.

[0085] The same survey further revealed the following clones with elevated expression levels in the presence of RA/ inhibitor:

55 HEMBA1000128, HEMBA1000875, HEMBA1001390, HEMBA1002163, HEMBA1002227, HEMBA1002421, HEMBA1004391, HEMBA1004454, HEMBA1004785, HEMBA1005913, HEMBA1006171, HEMBA1006299, HEMBA1006335, HEMBA1006544, HEMBA1007241, HEMBB1000447, HEMBB1000668, MAMMA1000994, MAMMA1001344, NT2RM2000582, NT2RP1001004, NT2RP2000663, NT2RP2000694, NT2RP2000903, NT2RP2001388, NT2RP2002674, NT2RP2002974, NT2RP2003383, NT2RP2004069, NT2RP2004606, NT2RP2004837, NT2RP2005069, NT2RP2005425, NT2RP2005463, NT2RP2005541, NT2RP2005883, NT2RP2005887, NT2RP3000460, NT2RP3000838, NT2RP3001044, NT2RP3001240, NT2RP3001388,

NT2RP3002721, NT2RP3002738, NT2RP3003469, NT2RP3004083, NT2RP3004130, NT2RP3004202,  
 NT2RP3004294, NT2RP3004640, NT2RP4000108, NT2RP4002451, NT2RP4002715, OVARC1000275,  
 OVARC1000467, OVARC1000553, OVARC1000853, OVARC1000873, OVARC1000916, OVARC1000995,  
 OVARC1001030, OVARC1001222, OVARC1001596, OVARC1002058, OVARC1002178, PLACE1000927,  
 5 PLACE1001123, PLACE1001407, PLACE1001464, PLACE1001564, PLACE1001795, PLACE1002547,  
 PLACE1003407, PLACE1003644, PLACE1003845, PLACE1004441, PLACE1004482, PLACE1005410,  
 PLACE1005601, PLACE1005725, PLACE1005736, PLACE1006093, PLACE1006219, PLACE1006290,  
 PLACE1006716, PLACE1007296, PLACE1007626, PLACE1008359, PLACE1010968, PLACE1011364,  
 PLACE1011824, THYRO1000678, THYRO1000776, THYRO1000999, THYRO1001113, THYRO1001237,  
 10 THYRO1001523, Y79AA1000226, Y79AA1000888, Y79AA1001430

**[0086]** The same survey further revealed the following clones with elevated expression levels in the presence of RA/ inhibitor:

HEMBA1000349, HEMBA1001297, HEMBA1001878, HEMBA1005070, HEMBA1006482, HEMBB1001959,  
 NT2RM2001939, NT2RP1000981, NT2RP2001469, NT2RP3003473, OVARC1001132, PLACE1001655,  
 15 Y79AA1000127, Y79AA1002381. These clones are associated with neural differentiation and, therefore, are candidates for genes associated with neurological diseases.

**[0087]** Based on the functional analyses using a secretory protein, membrane protein, or proteins associated with signal transduction, glycoprotein, transcription, or diseases, it is possible to develop a medicine.

**[0088]** In case of a membrane protein, it is most likely to be a protein that functions as a receptor or ligand on the cell surface. Therefore, it is possible to reveal a new relationship between a ligand and receptor by screening the membrane protein of the invention based on the binding activity with the known ligand or receptor. Screening can be performed according to the known methods.

**[0089]** For example, a ligand against the protein of the invention can be screened in the following manner. Namely, a ligand that binds to a specific protein can be screened by a method comprising the steps of: (a) contacting a test sample with the protein of the invention or a partial peptide thereof, or cells expressing these, and (b) selecting a test sample that binds to said protein, said partial peptide, or said cells.

**[0090]** On the other hand, for example, screening using cells expressing the protein of the present invention that is a receptor protein can also be performed as follows. It is possible to screen receptors that is capable of binding to a specific protein by using procedures (a) attaching the sample cells to the protein of the invention or its partial peptide, and (b) selecting cells that can bind to the said protein or its partial peptide.

**[0091]** In a following screening as an example, first the protein of the invention is expressed, and the recombinant protein is purified. Next, the purified protein is labeled, binding assay is performed using a various cell lines or primary cultured cells, and cells that are expressing a receptor are selected (Growth and differentiation factors and their receptors, Shin-Seikagaku Jikken Kouza Vol.7 (1991) Honjyo, Arai, Taniguchi, and Muramatsu edit, p203-236, Tokyo-Kagaku-Doujin). A protein of the invention can be labeled with RI such as <sup>125</sup>I, and enzyme (alkaline phosphatase etc.). Alternatively, a protein of the invention may be used without labeling and then detected by using a labeled antibody against the protein. The cells that are selected by the above screening methods, which express a receptor of the protein of the invention, can be used for the further screening of an agonists or antagonists of the said receptor.

**[0092]** Once the ligand binding to the protein of the invention, the receptor of the protein of the invention or the cells expressing the receptor are obtained by screening, it is possible to screen a compound that binds to the ligand and receptor. Also it is possible to screen a compound that can inhibit both bindings (agonists or antagonists of the receptor, for example) by utilizing the binding activities.

**[0093]** When the protein of the invention is a receptor, the screening method comprises the steps of (a) contacting the protein of the invention or cells expressing the protein of the invention with the ligand, in the presence of a test sample, (b) detecting the binding activity between said protein or cells expressing said protein and the ligand, and (c) selecting a compound that reduces said binding activity when compared to the activity in the absence of the test sample. Furthermore, when the protein of the invention is a ligand, the screening method comprises the steps of (a) contacting the protein of the invention with its receptor or cells expressing the receptor in the presence of samples, (b) detecting the binding activity between the protein and its receptor or the cells expressing the receptor, and (c) selecting a compound that can potentially reduce the binding activity compared to the activity in the absence of the sample.

**[0094]** Samples to screen include cell extracts, expressed products from a gene library, synthesized low molecular compound, synthesized peptide, and natural compounds, for example, but are not construed to be listed here. A compound that is isolated by the above screening using a binding activity of the protein of the invention can also be used as a sample.

**[0095]** A compound isolated by the screening may be a candidate to be an agonist or an antagonist of the receptor of the protein. By utilizing an assay that monitors a change in the intracellular signaling such as phosphorylation which results from reduction of the binding between the protein and its receptor, it is possible to identify whether the obtained compound is an agonist or antagonist of the receptor. Also, the compound may be a candidate of a molecule that can

inhibit the interaction between the protein and its associated proteins (including a receptor) in vivo. Such compounds can be used for developing drugs for precaution or cures of a disease with which the protein is associated.

[0096] Secretory proteins may regulate cellular conditions such as growth and differentiation. It is possible to find out a novel factor that regulates cellular conditions by adding the secretory protein of the invention to a certain kind of cell, and performing a screening by utilizing the cellular changes in growth or differentiation, or activation of a particular gene.

[0097] The screening can be performed, for example, as follows. First, the protein of the invention is expressed and purified in a recombinant form. Then, the purified protein is microinjected into a various kind of cell lines or primary cultured cells, and the change in the cell growth and differentiation is monitored. The induction of a particular gene that is known to be involved in a certain cellular change is detected with the amounts of mRNA and protein. Alternatively, the amount of an intracellular molecule (low molecular compounds, etc.) that is changed by the function of a gene product (protein) that is known to be functioning in a certain cellular change is used for the detection.

[0098] Once the screening reveals that the protein of the invention can regulate cellular conditions or the functions, it is possible to apply the protein as a pharmaceutical and diagnostic medicine for associated diseases by itself or by altering a part of it into an appropriate composition.

[0099] As is above described for membrane proteins, the secretory protein provided by the invention may be used to explore a novel ligand-receptor interaction using a screening based on the binding activity to a known ligand or receptor. A similar method can be used to identify an agonist or antagonist. The resulting compounds obtained by the methods can be a candidate of a compound that can inhibit the interaction between the protein of the invention and an interacting molecule (including a receptor). The compounds may be able to use as a preventive, therapeutic, and diagnostic medicine for the diseases, in which the protein may play a certain role.

[0100] Proteins associated with signal transduction or transcription may be a factor that affects a certain protein or gene in response to intracellular/extracellular stimuli. It is possible to find out a novel factor that can affect a protein or gene by expressing the protein provided by the invention in a certain types of cells, and performing a screening utilizing the activation of a certain intracellular protein or gene.

[0101] The screening may be performed as follows. First, a transformed cell expressing the protein is obtained. Then, the transformed cell line and the untransformed original cell are compared for the changes in the expression of a certain gene by detecting the amount of its mRNA or protein. Alternatively, the amount of an intracellular molecule (low molecular compounds), which is changed by the function of a gene product (protein) that is known to function in a certain cellular change, may be used for the detection. Furthermore, the change of the expression of a certain gene can be detected by introducing a fusion gene that comprises a regulatory region of the gene and a marker gene (luciferase, beta-galactosidase, etc.) into a cell, expressing the protein provided by the invention into the cell, and estimating the activity of a marker gene product (protein).

[0102] If the protein or gene of the invention is associated with diseases, it is possible to screen a gene or compound that can regulate its expression and/or activity either directly or indirectly by utilizing the protein of the present invention.

[0103] For example, the protein of the invention is expressed and the recombinant protein is purified. Then, the protein and gene whose expression was affected in the presence of the protein of the invention is also purified, and the binding activity between the two proteins or genes is examined. The examination may be performed with pretreatment with a compound that is candidate of an inhibitor. In an alternative method, a transcription regulatory region locating in the 5'-upstream of the gene encoding the protein of the invention that is capable of regulating the expression of other genes is obtained, and fused with a marker gene. The fusion is introduced into a cell, and the cell is added with compounds to explore a regulatory factor of the expression of the said gene.

[0104] The compound obtained by the screening can be used for developing pharmaceutical and diagnostic medicines for the diseases with which the protein of the present invention is associated. Similarly, if the regulatory factor obtained by the screening is a protein, the protein itself can be used as a pharmaceutical, and if there is a compound that affects the original expression level and/or activity of the protein, it also can be used for the same purpose.

[0105] If the protein of the invention has an enzymatic activity, regardless of whether it is a secretory protein, membrane protein, or proteins associated with signal transduction, glycoprotein, transcription, or diseases, a screening may be performed by adding a compound to the protein of the invention under the suitable condition and monitoring the change of the compound. The enzymatic activity may also be utilized to screen for a compound that can inhibit the activity of the protein.

[0106] In a screening given as an example, the protein of the invention is expressed and the recombinant protein is purified. Then, compounds are contacted with the purified protein, and the amount of the compound and the reaction products is examined. Alternatively, compounds that are candidates of an inhibitor are pretreated, then a compound (substrate) that can react with the purified protein is added, and the amount of the substrate and the reaction products is examined.

[0107] The compounds obtained in the screening may be used as a medicine for diseases with which the protein of the invention is associated. Also they can be applied for tests that examine whether the protein of the invention functions

normally *in vivo*.

**[0108]** Whether the secretory or membrane protein of the present invention is a novel protein associated with diseases or not is determined in another method than described above, by obtaining a specific antibody against the protein of the invention, and examining the relationship between the expression or activity of the protein and a certain disease. In an alternative way, it may be analyzed referred to the methods in "Molecular Diagnosis of Genetic Diseases" (Elles R. edit, (1996) in the series of "Method in Molecular Biology" (Humana Press).

**[0109]** Disease-associated proteins are a target of the above described screenings and very useful for developing a drug that is capable of regulating the expression and activity of the protein. Also, they are useful in medicinal industry as a diagnostic marker of the related disease and as a target for gene therapy.

**[0110]** Compounds isolated as mentioned above can be administered patients as it is, or after formulated into a pharmaceutical composition according to the known methods. For example, a pharmaceutically acceptable carrier or vehicle, specifically sterilized water, saline, plant oil, emulsifier, or suspending agent can be mixed with the compounds appropriately. The pharmaceutical compositions can be administered to patients by a method known to those skilled in the art, such as intraarterial intravenous, or subcutaneous injections. The dosage may vary depending on the weight or age of a patient, or the method of administration, but those skilled in the art can choose an appropriate dosage properly. If the compound is encoded by DNA, the DNA can be cloned into a vector for gene therapy, and used for gene therapy. The dosage of the DNA and the method of its administration may vary depending on the weight or age of a patient, or the symptoms, but those skilled in the art can choose properly.

**[0111]** The protein encoded by the polynucleotide of the invention can be prepared as a recombinant protein or as a natural protein. For example, the recombinant protein can be prepared by inserting the polynucleotide encoding the protein of the invention into a vector, introducing the vector into an appropriate host cell and purifying the protein expressed within the transformed host cell, as described below. In contrast, the natural protein can be prepared, for example, by utilizing an affinity column to which an antibody against the protein of the invention (Current Protocols in Molecular Biology (1987) Ausubel et al. edit, John Wiley & Sons, Section 16.1-16.19) is attached. The antibody used for the affinity purification may be either a polyclonal antibody, or a monoclonal antibody. Alternatively, *in vitro* translation (See, for example, "On the fidelity of mRNA translation in the nuclease-treated rabbit reticulocyte lysate system." Dasso M.C., and Jackson R.J. (1989) Nucleic Acids Res. 17: 3129-3144) may be used for preparing the protein of the invention.

**[0112]** Proteins functionally equivalent to the proteins of the present invention can be prepared based on the activities, which were clarified in the above-mentioned manner, of the proteins of the present invention. Using the biological activity possessed by the protein of the invention as an index, it is possible to verify whether or not a particular protein is functionally equivalent to the protein of the invention by examining whether or not the protein has said activity.

**[0113]** Proteins functionally equivalent to the proteins of the present invention can be prepared by those skilled in the art, for example, by using a method for introducing mutations into an amino acid sequence of a protein (for example, site-directed mutagenesis (Current Protocols in Molecular Biology, edit, Ausubel et al., (1987) John Wiley & Sons, Section 8.1-8.5). Besides, such proteins can be generated by spontaneous mutations. The present invention comprises the proteins having one or more amino acid substitutions, deletions, insertions and/or additions in the amino acid sequences of the proteins of the present invention (Table 370), as far as the proteins have the equivalent functions to those of the proteins identified in the present Examples described later.

**[0114]** There are no limitations in the number and sites of amino acid mutations, as far as the proteins maintain the functions thereof. The number of mutations is typically 30% or less, or 20% or less, or 10% or less, preferably within 5% or less, or 3% or less of the total amino acids, more preferably within 2% or less or 1% or less of the total amino acids. From the viewpoint of maintaining the protein function, it is preferable that a substituted amino has a similar property to that of the original amino acid. For example, Ala, Val, Leu, Ile, Pro, Met, Phe and Trp are assumed to have similar properties to one another because they are all classified into a group of non-polar amino acids. Similarly, substitution can be performed among non-charged amino acid such as Gly, Ser, Thr, Cys, Tyr, Asn, and Gln, acidic amino acids such as Asp and Glu, and basic amino acids such as Lys, Arg, and His.

**[0115]** In addition, proteins functionally equivalent to the proteins of the present invention can be isolated by using techniques of hybridization or gene amplification known to those skilled in the art. Specifically, using the hybridization technique (Current Protocols in Molecular Biology, edit, Ausubel et al., (1987) John Wiley & Sons, Section 6.3-6.4)), those skilled in the art can usually isolate a DNA highly homologous to the DNA encoding the protein identified in the present Example based on the identified nucleotide sequence (Table 370) or a portion thereof and obtain the functionally equivalent protein from the isolated DNA. The present invention includes proteins encoded by the DNAs hybridizing with the DNAs encoding the proteins identified in the present Example, as far as the proteins are functionally equivalent to the proteins identified in the present Example. Organisms from which the functionally equivalent proteins are isolated are illustrated by vertebrates such as human, mouse, rat, rabbit, pig and bovine, but are not limited to these animals.

**[0116]** Washing conditions of hybridization for the isolation of DNAs encoding the functionally equivalent proteins are usually "1xSSC, 0.1% SDS, 37°"; more stringent conditions are "0.5xSSC, 0.1% SDS, 42°"; and still more stringent conditions are "0.1 x SSC, 0.1% SDS, 65°". Alternatively, the following conditions can be given as hybridization con-

ditions of the present invention. Namely, conditions in which the hybridization is done at "6xSSC, 40% Formamide, 25.", and the washing at "1xSSC, 55." can be given. More preferable conditions are those in which the hybridization is done at "6xSSC, 40% Formamide, 37.", and the washing at "0.2xSSC, 55.". Even more preferable are those in which the hybridization is done at "6xSSC, 50% Formamide, 37.", and the washing at "0.1xSSC, 62.". The more stringent  
 5 the conditions of hybridization are, the more frequently the DNAs highly homologous to the probe sequence are isolated. Therefore, it is preferable to conduct hybridization under stringent conditions. Examples of stringent conditions in the present invention are, washing conditions of "0.5xSSC, 0.1% SDS, 42.", or alternatively, hybridization conditions of "6xSSC, 40% Formamide, 37.", and the washing at "0.2xSSC, 55.". However, the above-mentioned combinations of  
 10 SSC, SDS and temperature conditions are indicated just as examples. Those skilled in the art can select the hybridization conditions with similar stringency to those mentioned above by properly combining the above-mentioned or other factors (for example, probe concentration, probe length and duration of hybridization reaction) that determines the stringency of hybridization.

**[0117]** The amino acid sequences of proteins isolated by using the hybridization techniques usually exhibit high homology to those of the proteins of the present invention, which are shown in Table 370. The present invention encompasses a polynucleotide comprising a nucleotide sequence that has a high identity to the nucleotide sequence of claim 8 (a). Furthermore, the present invention encompasses a peptide, or protein comprising an amino acid sequence that has a high identity to the amino acid sequence encoded by the polynucleotide of claim 8 (b). The term "high identity" indicates sequence identity of at least 40% or more; preferably 60% or more; and more preferably 70% or more. Alternatively, more preferable is identity of 90% or more, or 93% or more, or 95% or more, furthermore, 97% or more, or 99% or more. The identity can be determined by using the BLAST search algorithm.

**[0118]** With the gene amplification technique (PCR) (Current Protocols in Molecular Biology, edit, Ausubel et al., (1987) John Wiley & Sons, Section 6.3-6.4)) using primers designed based on the nucleotide sequence (Table 370) or a portion thereof identified in the present Example, it is possible to isolate a DNA fragment highly homologous to the nucleotide sequence or a portion thereof and to obtain functionally equivalent protein to a particular protein identified  
 25 in the present Example based on the isolated DNA fragment.

**[0119]** The "percent identity" of two amino acid sequences or of two nucleic acids is determined using the algorithm of Karlin and Altschul (Proc. Natl. Acad. Sci. USA 87:2264-2268, 1990), modified as in Karlin and Altschul (Proc. Natl. Acad. Sci. USA 90:5873-5877, 1993). Such an algorithm is incorporated into the BLASTN and BLASTX programs of Altschul et al. (J. Mol. Biol. 215:403-410, 1990). BLAST nucleotide searches are performed with the BLASTN program, score = 100, wordlength = 12. BLAST protein searches are performed with the BLASTX program, score = 50, wordlength = 3. When gaps exist between two sequences, Gapped BLAST is utilized as described in Altschul et al. (Nucleic Acids Res. 25:3389-3402, 1997). When utilizing BLAST and Gapped BLAST programs, the default parameters of the respective programs (e.g., BLASTX and BLASTN) are used. See <http://www.ncbi.nlm.nih.gov>.

**[0120]** The present invention also includes a partial peptide of the proteins of the invention. The partial peptide comprises a protein generated as a result that a signal peptide has been removed from a secretory protein. If the protein of the present invention has an activity as a receptor or a ligand, the partial peptide may function as a competitive inhibitor of the protein and may bind to the receptor (or ligand). In addition, the present invention comprises an antigen peptide for raising antibodies. For the peptides to be specific for the protein of the invention, the peptides comprise at least 7 amino acids, preferably 8 amino acids or more, more preferably 9 amino acids or more, and even more preferably  
 40 10 amino acids or more. The peptide can be used for preparing antibodies against the protein of the invention, or competitive inhibitors of them, and also screening for a receptor that binds to the protein of the invention. The partial peptides of the invention can be produced, for example, by genetic engineering methods, known methods for synthesizing peptides, or digesting the protein of the invention with an appropriate peptidase.

**[0121]** The present invention also relates to a vector into which the DNA of the invention is inserted. The vector of the invention is not limited as long as it contains the inserted DNA stably. For example, if E. coli is used as a host, vectors such as pBluescript vector (Stratagene) are preferable as a cloning vector. To produce the protein of the invention, expression vectors are especially useful. Any expression vector can be used as far as it is capable of expressing the protein in vitro, in E. coli, in cultured cells, or in vivo. For example, pBEST vector (Promega) is preferable for in vitro expression, pET vector (Invitrogen) for E. coli, pME18S-FL3 vector (GenBank Accession No. AB009864) for cultured cells, and pME18S vector (Mol. Cell. Biol. (1988) 8: 466-472) for in vivo expression. To insert the DNA of the invention, ligation utilizing restriction sites can be performed according to the standard method (Current Protocols in Molecular Biology (1987) Ausubel et al. edit, John Wiley & Sons, Section 11.4-11.11).

**[0122]** The present invention also relates to a transformant carrying the vector of the invention. Any cell can be used as a host into which the vector of the invention is inserted, and various kinds of host cells can be used depending on the purposes. For strong expression of the protein in eukaryotic cells, COS cells or CHO cells can be used, for example.

**[0123]** Introduction of the vector into host cells can be performed, for example, by calcium phosphate precipitation method, electroporation method (Current Protocols in Molecular Biology (1987) Ausubel et al. edit, John Wiley & Sons, Section 9.1-9.9), lipofectamine method (GIBCO-BRL), or microinjection method, etc.



**[0124]** The primer of the present invention can be used for synthesizing full-length cDNA, and also for the detection and/or diagnosis of the abnormality of the protein of the invention encoded by the full-length cDNA. For example, by utilizing polymerase chain reaction (genomic DNA-PCR, or RT-PCR) using the primer of the invention, DNA encoding the protein of the invention can be amplified. It is also possible to obtain the regulatory region of expression in the 5'-upstream by using PCR or hybridization since the transcription start site within the genomic sequence can be easily specified based on the 5'-end sequence of the full-length cDNA. The obtained genomic region can be used for detection and/or diagnosis of the abnormality of the sequence by RFLP analysis, SSCP, or direct sequencing.

**[0125]** Furthermore, the "polynucleotide having a length of at least 15 nucleotides, comprising a nucleotide sequence that is complementary to a polynucleotide comprising the nucleotide sequence set forth in any one of SEQ ID NOs in Table 370, or its complementary strand" includes an antisense polynucleotide for suppressing the expression of the protein of the invention. To exert the antisense effect, the antisense polynucleotide has a length of at least 15 bp or more, for example, 50 bp or more, preferably 100 bp or more, and more preferably 500 bp or more, and has a length of usually 3000 bp or less and preferably 2000 bp or less. The antisense DNA can be used in the gene therapy of the diseases that are caused by the abnormality of the protein of the invention (abnormal function or abnormal expression). Said antisense DNA can be prepared, for example, by the phosphorothioate method ("Physicochemical properties of phosphorothioate oligodeoxynucleotides." Stein (1988) *Nucleic Acids Res.* 16: 3209-3221) based on the nucleotide sequence of the DNA encoding the protein (for example, the DNA set forth in any one of SEQ ID NOs in Table 370).

**[0126]** The polynucleotide or antisense DNA of the present invention can be used in gene therapy, for example, by administering it into a patient by the in vivo or ex vivo method with virus vectors such as retrovirus vectors, adenovirus vectors, and adeno-associated virus vectors, or non-virus vectors such as liposome.

**[0127]** The present invention also relates to antibodies that bind to the protein of the invention. There are no limitations in the form of the antibodies of the invention. They include polyclonal antibodies, monoclonal antibodies, or their portions that can bind to the protein of the invention. They also include antibodies of all classes. Furthermore, special antibodies such as humanized antibodies are also included.

**[0128]** The polyclonal antibody of the invention can be obtained according to the standard method by synthesizing an oligopeptide corresponding to the amino acid sequence and immunizing rabbits with the peptide (Current Protocols in Molecular Biology (1987) Ausubel et al. edit, John Wiley & Sons, Section 11.12-11.13). The monoclonal antibody of the invention can be obtained according to the standard method by purifying the protein expressed in *E. coli*, immunizing mice with the protein, and producing a hybridoma cell by fusing the spleen cells and myeloma cells (Current Protocols in Molecular Biology (1987) Ausubel et al. edit, John Wiley & Sons, Section 11.4-11.11).

**[0129]** The antibody binding to the protein of the present invention can be used for purification of the protein of the invention, and also for detection and/or diagnosis of the abnormalities of the expression and structure of the protein. Specifically, proteins can be extracted, for example, from tissues, blood, or cells, and the protein of the invention is detected by Western blotting, immunoprecipitation, or ELISA, etc. for the above purpose.

**[0130]** Furthermore, the antibody binding to the protein of the present invention can be utilized for treating the diseases that associates with the protein of the invention. If the antibodies are used for treating patients, human antibodies or humanized antibodies are preferable in terms of their low antigenicity. The human antibodies can be prepared by immunizing a mouse whose immune system is replaced with that of human ("Functional transplant of megabase human immunoglobulin loci recapitulates human antibody response in mice" Mendez M.J. et al. (1997) *Nat. Genet.* 15: 146-156). The humanized antibodies can be prepared by recombination of the hypervariable region of a monoclonal antibody (Methods in Enzymology (1991) 203: 99-121).

**[0131]** The present invention further relates to databases comprising at least a sequence of polynucleotide and/or protein, or a medium recorded in such databases, selected from the sequence data of the nucleotide and/or the amino acids indicated in Table 370. The term "database" means a set of accumulated information as machine-searchable and readable information of nucleotide sequence. The databases of the present invention comprise at least one of the novel nucleotide sequences of polynucleotide provided by the present invention. The databases of the present invention can consist of only the sequence data of the polynucleotide provided by the present invention or can comprise other information on nucleotide sequences of known full-length cDNAs or ESTs. The databases of the present invention can be comprised of not only the information on the nucleotide sequences but also the information on the gene functions revealed by the present invention. Additional information such as names of DNA clones carrying the full-length cDNAs can be recorded or linked together with the sequence data in the databases.

**[0132]** The database of the present invention is useful for gaining complete gene sequence information from partial sequence information of a gene of interest. The database of the present invention comprises nucleotide sequence information of full-length cDNAs. Consequently, by comparing the information in this database with the nucleotide sequence of a partial gene fragment yielded by differential display method or subtraction method, the information on the full-length nucleotide sequence of interest can be gained from the sequence of the partial fragment as a starting clue.

**[0133]** The sequence information of the full-length cDNAs constituting the database of the present invention contains not only the information on the complete sequences but also extra information on expression frequency of the genes

as well as homology of the genes to known genes and known proteins. Thus the extra information facilitates rapid functional analyses of partial gene fragments. Further, the information on human genes is accumulated in the database of the present invention, and therefore, the database is useful for isolating a human homologue of a gene originating from other species. The human homologue can be isolated based on the nucleotide sequence of the gene from the original species.

**[0134]** At present, information on a wide variety of gene fragments can be obtained by differential display method and subtraction method. In general, these gene fragments are utilized as tools for isolating the full-length sequences thereof. When the gene fragment corresponds to an already-known gene, the full-length sequence is easily obtained by comparing the partial sequence with the information in known databases. However, when there exists no information corresponding to the partial sequence of interest in the known databases, cDNA cloning should be carried out for the full-length cDNA. It is often difficult to obtain the full-length nucleotide sequence using the partial sequence information as an initial clue. If the full-length of the gene is not available, the amino acid sequence of the protein encoded by the gene remains unidentified. Thus the database of the present invention can contribute to the identification of full-length cDNAs corresponding to gene fragments, which cannot be revealed by using databases of known genes.

**[0135]** The invention is illustrated more specifically with reference to the following examples, but is not to be construed as being limited thereto.

#### EXAMPLE 1

Construction of a cDNA library by the oligo-capping method.

**[0136]** The NT-2 neuron progenitor cells (Stratagene), a teratocarcinoma cell line from human embryo testis, which can differentiate into neurons by treatment with retinoic acid were used. The NT-2 cells were cultured according to the manufacturer's instructions as follows.

- (1) NT-2 cells were cultured without induction by retinoic acid treatment ((NT2RM1, NT2RM2, NT2RM4)).
- (2) After cultured, NT-2 cells were induced by adding retinoic acid, and then were cultured for 48 hours (NT2RP1).
- (3) After cultured, NT-2 cells were induced by adding retinoic acid, and then were cultured for 2 weeks (NT2RP2, NT2RP3, NT2RP4).

**[0137]** Also, the human brain neuroglioma cell line H4 (ATCC HTG-148) (BNGH41), human neuroblastoma cell line SK-N-MC (ATCC HTB-10) (SKNMC1), and human retinoblastoma cell line Y79 (ATCC HTB-18) (Y79AA1) were cultured according to the culture conditions described in the ATCC catalogue. The cells were harvested separately, and mRNA was extracted from each cell by the method described in the literature (Molecular Cloning 2nd edition. Sambrook J., Fritsch, E.F., and Maniatis T. (1989) Cold Spring Harbor Laboratory Press). Furthermore, poly(A)<sup>+</sup>RNA was purified from the mRNA using oligo-dT cellulose.

**[0138]** Similarly, human placenta (PLACE1, PLACE2, PLACE3), human ovary cancer tissue (OVARC1), tissues from human embryo at 10 weeks, which is enriched with head (HEMBA1), or body (HEMBB1), human mammary gland (MAMMA1), human thyroid gland (THYRO1) were used to extract mRNA by the method described in the literature (Molecular Cloning 2nd edition. Sambrook J., Fritsch, E.F., and Maniatis T. (1989) Cold Spring Harbor Laboratory Press). Furthermore, poly(A)<sup>+</sup>RNA was purified from the mRNA using oligo-dT cellulose.

**[0139]** Each poly(A)<sup>+</sup>RNA was used to construct a cDNA library by the oligo-capping method (Maruyama M. and Sugano S. (1994) Gene 138: 171-174). Using the Oligo-cap linker (SEQ ID NO: 2541) and the Oligo-dT primer (SEQ ID NO: 2542), bacterial alkaline phosphatase (BAP) treatment, tobacco acid phosphatase (TAP) treatment, RNA ligation, the first strand cDNA synthesis, and removal of RNA were performed as described in the reference (Suzuki and Kanno (1996) Protein Nucleic acid and Enzyme. 41: 197-201; Suzuki Y. et al. (1997) Gene 200: 149-156). Next, 5'- and 3'-PCR primers (SEQ ID NO: 2543, and 2544, respectively) were used for performing PCR to convert the cDNA into double stranded cDNA, which was then digested with SfiI. Then, the DraIII-cleaved pUC19FL3 vector (Figure 1; for NT2RM1, and NT2RP1), or the DraIII-cleaved pME18SFL3 (Figure 1) (GenBank AB009864, expression vector; for NT2RM2, NT2RM4, NT2RP2, NT2RP3, NT2RP4, BNGH41, SKNMC1, Y79AA1, PLACE1, PLACE2, PLACE3, OVARC1, HEMBA1, HEMBB1, MAMMA1, and THYRO1) was used for cloning the cDNA in an unidirectional manner, and cDNA libraries were obtained. Then, the nucleotide sequence of the 5'- and 3'- ends of the cDNA clones was analyzed with a DNA sequencer (ABI PRISM 377, PE Biosystems) after sequencing reactions were performed with the DNA sequencing reagents (Dye Terminator Cycle Sequencing FS Ready Reaction Kit, dRhodamine Terminator Cycle Sequencing FS Ready Reaction Kit, or BigDye Terminator Cycle Sequencing FS Ready Reaction Kit, from by PE Biosystems) according to the instructions. The data were compiled into a database.

**[0140]** The full-length-enriched cDNA libraries except those for NT2RM1 and NT2RP1 were constructed using eukaryotic expression vector pME18SFL3. The vector contains SR $\alpha$  promoter and SV40 small t intron in the upstream

of the cloning site, and SV40 polyA added signal sequence site in the downstream. As the cloning site of pME18SFL3 has asymmetrical *Dra*III sites, and the ends of cDNA fragments contain *Sfi*I sites complementary to the *Dra*III sites, the cloned cDNA fragments can be inserted into the downstream of the SR $\alpha$  promoter unidirectionally. Therefore, clones containing full-length cDNA can be expressed transiently by introducing the obtained plasmid directly into COS cells. Thus, the clones can be analyzed very easily in terms of the proteins that are the gene products of the clones, or in terms of the biological

[0141] Herein, the cDNA libraries and the name of each clone are related as shown in Table 2. Therein, "xxxxxx" represents the clone number of six digits. Thus, the sequences are named by the library name, the clone number plus F- for the 5'-end, or R- for the 3'-end.

Table 2

library:

	clone	5'-end sequence	3'-end sequence
5			
	NT2RM1:		
	NT2RM1xxxxxx	F-NT2RM1xxxxxx	
10	NT2RP1:		
	NT2RP1xxxxxx	F-NT2RP1xxxxxx	
	NT2RM2:		
	NT2RM2xxxxxx	F-NT2RM2xxxxxx	R-NT2RM2xxxxxx
15	NT2RM4:		
	NT2RM4xxxxxx	F-NT2RM4xxxxxx	R-NT2RM4xxxxxx
	NT2RP2:		
20	NT2RP2xxxxxx	F-NT2RP2xxxxxx	R-NT2RP2xxxxxx
	NT2RP3:		
	NT2RP3xxxxxx	F-NT2RP3xxxxxx	R-NT2RP3xxxxxx
	NT2RP4:		
25	NT2RP4xxxxxx	F-NT2RP4xxxxxx	R-NT2RP4xxxxxx
	BNGH41:		
	BNGH41xxxxxx	F-BNGH41xxxxxx	R-BNGH41xxxxxx
30	SKNMC1:		
	SKNMC1xxxxxx	F-SKNMC1xxxxxx	R-SKNMC1xxxxxx
	Y79AA1:		
	Y79AA1xxxxxx	F-Y79AA1xxxxxx	R-Y79AA1xxxxxx
35	PLACE1:		
	PLACE1xxxxxx	F-PLACE1xxxxxx	R-PLACE1xxxxxx
	PLACE2:		
40	PLACE2xxxxxx	F-PLACE2xxxxxx	R-PLACE2xxxxxx
	PLACE3:		
	PLACE3xxxxxx	F-PLACE3xxxxxx	R-PLACE3xxxxxx
	OVARC1:		
45	OVARC1xxxxxx	F-OVARC1xxxxxx	R-OVARC1xxxxxx
	HEMBA1:		
	HEMBA1xxxxxx	F-HEMBA1xxxxxx	R-HEMBA1xxxxxx
50	HEMBB1:		
	HEMBB1xxxxxx	F-HEMBB1xxxxxx	R-HEMBB1xxxxxx
	MAMMA1:		
	MAMMA1xxxxxx	F-MAMMA1xxxxxx	R-MAMMA1xxxxxx
55	THYRO1:		

THYRO1xxxxxx F-THYRO1xxxxxx R-THYRO1xxxxxx

5

**EXAMPLE 2**

Estimation of the fullness ratio of the 5'-ends of the clones contained in the cDNA libraries constructed by the oligo-capping method.

10

[0142] The fullness ratio at the 5'-end sequences of the 59,823 clones in the human cDNA libraries constructed by the oligo-capping method was determined as follows. Of all the clones whose 5'-end sequences were found in those of known human mRNA in the public database, a clone was judged to be "full-length", if it had a longer 5'-end sequence than that of the known human mRNA, or, even though the 5'-end sequence was shorter, if it contained the translation initiation codon. A clone which did not contain the translation initiation codon was judged to be "non-full-length". The fullness ratio ((the number of full-length clones)/(the number of full-length and non-full-length clones)) at the 5'-end of the cDNA clones from each library was determined by comparing with the known human mRNA. As a result, the fullness ratio of the 5'-ends was 63.5%. It suggests that the human cDNA clones obtained by the described method have complete 5'-ends with high probability.

20

**EXAMPLE 3**

Assessment of the fullness ratio of the 5'-end of the cDNA by the ATGpr and the ESTimateFL.

25

[0143] The ATGpr, developed by Salamov A.A., Nishikawa T., and Swindells M.B. in the Helix Research Institute, is a program for prediction of the translation start codon based on the characteristics of the sequences in the vicinity of the ATG codon. The results are shown with expectations that an ATG is a true start codon (0.05-0.94). When this program is applied to general cDNAs without considering whether or not the ATG codons in the cDNAs are the true initiation codons of the cDNAs, both the sensitivity and the specificity of the results are estimated at 66%. Here, the sensitivity means the ratio of the number of codons judged to be initiation codons by the program to the total number of true initiation codons, and the specificity means the ratio of the number of true initiation codons to the number of codons judged to be initiation codons by the program. In contrast, when the program was applied to the 5'-end sequences of the clones from the cDNA library that was obtained by the oligo-capping method and that had 65% fullness ratio, the sensitivity and specificity of evaluation of a full-length clone (clone containing the N-terminal end of ORF) were improved to 82-83% by selecting only clones having the ATGpr1 score 0.6 or higher.

35

[0144] Furthermore, the program was used to assess the fullness of 18,959 clones in the human cDNA libraries obtained here, which have 5'-ends matched to a known human mRNA. Briefly, the maximal ATGpr1 score of the clones was determined, and then their 5'-end sequence was compared with the known human mRNA to estimate whether the clone is full-length or not. The result was summarized in Table 3. Based on the knowledge that known mRNAs, in general, are highly expressed in the cell, the expression levels of genes having a low number in the EST hit, which represent mRNAs whose expression levels are relatively low were examined, and the result is shown in Table 4.

40

[0145] In the table, the number of full-length clones indicate that of clones containing the N-terminal end of ORF, and so does the number of non-full-length clones that of clones without the N-terminal end of ORF. The fullness ratio represents (the number of full-length clones)/(the number of full-length clones plus the number of non-full-length clones).

45

Table 3

The maximal ATGpr1 score and the fullness ratio of the 5'-end sequences of clones obtained from human cDNA libraries constructed by the oligo-capping method; clones having a matched 5'-end with that of a known human mRNA.

50

maximal ATGpr1 score	number of (full-length clones plus non-full-length clones)	number of full-length clones	fullness ratio
>=0.70	11,193	9,346	83.5%
>=0.50	13,369	10,549	78.9%
>=0.30	15,489	11,340	73.2%

55

Table 3 (continued)

The maximal ATGpr1 score and the fullness ratio of the 5'-end sequences of clones obtained from human cDNA libraries constructed by the oligo-capping method; clones having a matched 5'-end with that of a known human mRNA.			
maximal ATGpr1 score	number of (full-length clones plus non-full-length clones)	number of full-length clones	fullness ratio
$\geq 0.15$	17,394	11,811	67.9%
$\geq 0.00$	18,959	12,046	63.5%

Table 4

The maximal ATGpr1 score and the fullness ratio of the 5'-end sequences of the clones obtained from human cDNA libraries constructed by the oligo-capping method; clones having 5 EST hits or less among the clones having a matched 5'-end with that of a known human mRNA.			
maximal ATGpr1 score	number of (full-length clones plus non-full-length clones)	number of full-length clones	fullness ratio
$\geq 0.70$	2,801	1,934	69.0%
$\geq 0.50$	3,683	2,393	65.0%
$\geq 0.30$	4,683	2,707	57.8%
$\geq 0.15$	5,559	2,890	52.0%
$\geq 0.00$	6,113	3,013	49.8%

[0146] The ESTiMateFL, developed by Nishikawa and Ota in the Helix Research Institute, is a method for the selection of a clone with high fullness ratio by comparing with the 5'-end or 3'-end sequences of ESTs in the public database.

[0147] By the method, a cDNA clone is judged presumably not to be full-length if there exist any ESTs which have longer 5'-end or 3'-end sequences than the clone. The method is systematized for high throughput analysis. A clone is judged to be full-length if the clone has a longer 5'-end sequence than ESTs in the public database. Even if a clone has a shorter 5'-end, the clone is judged to be full-length if the difference in length is within 50 bases, and otherwise judged not to be full-length, for convenience. In case of the 5'-end sequence of the clones which matches a known mRNA, about 80% of the sequences that were judged to be full-length by comparing with ESTs was judged to be full-length by estimating the 5'-end sequence, as well; about 80% of the sequences that were judged to be not full-length by comparing with ESTs was judged to be not full-length by estimating the 5'-end sequence, as well. The accuracy of the prediction by comparing cDNA clones with ESTs is improved with increasing number of ESTs to be compared. However, when only a limited number of ESTs are available, the reliability becomes low. Thus, the method is effective in excluding clones with high probability of being non-full-length, from the cDNA clones that is synthesized by the oligo-capping method and that have the 5'-end sequences with about 60 % fullness ratio. In particular, the ESTiMateFL is efficiently used to estimate the fullness ratio at the 3'-end sequence of cDNA of a human unknown mRNA which has a significant number of ESTs in the public database.

[0148] The 18,959 clones isolated from human cDNA libraries constructed by the oligo-capping method, which have the 5'-end sequence that matches a known human mRNA, were estimated by using the ATGpr and ESTiMateFL. Briefly, the 5'-end sequence that matches a known human mRNA of the respective clone was analyzed to obtain the maximal ATGpr1 score, and compared with the ORF of the known human mRNA that matches it to determine whether the clone is full-length or not. Then, the 5'-end sequence of the respective clone was analyzed by the ESTiMateFL to judge whether the clone is full-length or not. Specifically, the 5'-end sequences that match a known human mRNA of the 18,959 clones constructed by the oligo-capping method were compared with those of ESTs by the ESTiMateFL and the clones other than those that are not full-length were selected. Then, the selected clones were used to analyze the relationship between the ATGpr and the fullness ratio. The result was summarized in Table 5. Also, among the selected, the clones in which the number of the EST hit is not more than 5 were selected and analyzed. The result was summarized in Table 6, which represents the result of the analysis of mRNA with relatively low abundance.

[0149] In the Tables, the number of full-length clones, the number of non-full-length clones, and the fullness ratio indicate the number of the clones that contain the N-terminus of the ORF, the number of the clones that do not contain

the N-terminus of the ORF, and (the number of full-length clones)/(the number of full-length clones plus (the number of non-full-length clones), respectively.

Table 5

The maximal ATGpr1 score and the fullness ratio of the 5'-end sequence in the clones isolated from human cDNA libraries constructed by the oligo-capping method, which have the 5'-end sequence that matches a known human mRNA, and also other than those being not full-length according to the comparison with ESTs.			
maximal ATGpr1 score	number of (full-length clones plus non-full-length clones)	number of full-length clones	fullness ratio
>=0.70	9,068	8,349	92.1%
>=0.50	10,345	9,318	90.1%
>=0.30	11,425	9,964	87.2%
>=0.15	12,254	10,335	84.3%
>=0.00	12,785	10,484	82.0%

Table 6

The maximal ATGpr1 score and the fullness ratio of the 5'-end sequence in the clones isolated from human cDNA libraries constructed by the oligo-capping method, which have the 5'-end sequence that matches a known human mRNA, and also other than those being not full-length according to the comparison with ESTs, in which the number of the EST hit is not more than 5.			
maximal ATGpr1 score	number of (full-length clones plus non-full-length clones)	number of full-length clones	fullness ratio
>=0.70	1,959	1,510	77.1%
>=0.50	2,469	1,821	73.8%
>=0.30	2,975	2,046	68.8%
>=0.15	3,368	2,164	64.3%
>=0.00	3,661	2,226	60.8%

[0150] According to the above results, it was found that, in case of using clones isolated from human cDNA libraries constructed by the oligo-capping method, the fullness ratio of the clones that have low score in the ATGpr can be improved by assessing their 5'-end sequence using the combination of the ATGpr and the ESTiMateFL. Therefore, the method was applied to select a cDNA clone with high fullness ratio.

#### EXAMPLE 4

Clustering of the 5'-end and 3'-end sequences of cDNA clones.

[0151] The 5'-end and 3'-end sequences of cDNA clones were obtained, and clustered separately. Briefly, data of the single pass sequencing of the determined 5'-end and 3'-end of cDNA clones was subjected to the BLAST search between the sequence data of all the clones synthesized in Example 1, and clones that are supposed to be originating from the same gene were clustered into a group. For the 5'-end sequence, those having the consensus sequence of 95% identity 300 base pairs or more are clustered into the same group. For the 3'-end sequence, those having the consensus sequence of 90% identity 200 base pairs or more are clustered into the same group. Among the clusters of the 5'-end and 3'-end sequences, the sequence having the longest lead was chosen as the representative sequence of the cluster (group).

#### EXAMPLE 5

Characterization of the representative sequences and the sequences of clones

[0152] Data of the 5'-end sequences of the representative sequences and clones was characterized by the following

methods:

(1) judging whether it is identical to the sequence of mRNA or ESTs from human by the BLAST search of the GenBank or SwissProt, and examining whether it is full-length by comparing with the sequences of known mRNA and ESTs from human.

(2) determining the ATGpr1 score using all the initiation codons contained within the 5'-end sequence by the ATGpr which predict fullness ratio.

(3) predicting the existence of the signal sequence using all the initiation codons contained within the 5'-end sequence by the PSORT which predict signal.

and,

(4) only with the 5'-end sequences of the representative sequences of the clusters, examining the keywords in the top hit data of the homology search of the SwissProt.

[0153] Data of the characterized representative sequences and clones was used for the final selection of the clones.

#### EXAMPLE 6

Identity to the human mRNA and human EST, and comparison of the 5'-end length.

[0154] The clones and the representative sequences of the clusters were judged to be identical to any human mRNA, if their 5'-end sequence has a region of 200 nucleotides or longer with 94% or more identity to the mRNA. The clones and the representative sequences of the clusters were judged to be identical to any human EST, if their 5'-end sequence has a region of 200 nucleotides or longer with 90% or more identity to the EST.

[0155] The clones and the representative sequences of the clusters were judged to be full-length in comparison with human mRNA, if their 5'-end sequence is longer than those of the mRNA, or it contains the translation initiation site. The clones and the representative sequences of the clusters were judged to be full-length in comparison with human EST in the database, if their 5'-end sequence is longer than those of the EST, or even though it is shorter, the difference in length between the two sequences is 50 nucleotides or less, for convenience. Otherwise, the clones and the representative sequences of the clusters were judged to be not full-length.

#### EXAMPLE 7

Prediction of the fullness ratio by the ATGpr.

[0156] The score in the ATGpr1 is the expectation to be full-length based on calculations, and the higher score reflects the higher fullness ratio as shown in Example 3. Further, the maximal ATGpr1 score represents the score obtained with all the initiation codons contained in the 5'-end sequence of the clones and the representative sequences, and are used for the characterization.

#### EXAMPLE 8

Prediction of the existence of a signal sequence by the PSORT.

[0157] Prediction of the existence of a signal sequence by the PSORT was performed on all of the amino acid sequences predicted from all the initiation codons in the 5'-end sequence of the clones and the representative sequences of the clusters. By analyzing the presence or absence of the sequence which is predicted to be a signal sequence, which is characteristics of the N-terminus of many secretory proteins, cDNA clones encoding a secretory protein or membrane protein were selected.

#### EXAMPLE 9

Prediction of the protein function by the BLAST search.

[0158] The 5'-end sequence of the representative sequences of the cluster was analyzed by the BLAST homology search of the SwissProt. The obtained top hit data was classified into those identical to the 5'-end representative sequence (identity was 90% or higher), those not identical to the 5'-end representative sequence (identity was 60% or lower, and compared sequence was not more than 25 nucleotides), and those similar to the 5'-end representative sequence (the rest of the data).



[0159] All the keywords in the SwissProt data corresponding to the top hit data were selected, and the 5'-end representative sequences were classified by the keywords relating with functions.

The keywords relating with a secretory protein or membrane protein are the followings:

5 growth factor,  
cytokine,  
hormone,  
receptor,  
G-protein coupled receptor,  
10 ionic channel,  
voltage-gated channel,  
calcium channel,  
extracellular matrix,  
transmembrane, and  
15 signal.

[0160] The keywords relating to glycoprotein is glycoprotein.

[0161] The keywords relating to signal transduction are the followings:

20 serine/threonine-protein kinase,  
tyrosine-protein kinase, and  
calmodulin-binding.

[0162] The keywords relating to transcription are the followings:

25 transcription regulation and activator,  
transcription regulation and repressor, and  
nuclear protein and repressor.

30 [0163] The keywords relating to diseases are disease mutation, and syndrome.

[0164] Many keywords overlapped in the respective group (receptor and transmembrane, for example), and some keywords overlapped in different groups (secretory or membrane, and diseases, etc.).

#### EXAMPLE 10

35 Selection of clones by characterization.

[0165] From the data obtained by the above characterization, clones encoding a novel secretory protein or membrane protein, or proteins with other predicted functions were selected by the combination of the ATGpr1 score and the prediction of the signal sequence by the PSORT, or according to the top hit data in the homology search of the SwissProt.

40 [0166] In selecting the clones, the 5'-end sequences that are identical to a human mRNA were ignored, whereas those that are identical to a human mRNA in part but obviously not identical in the other part were included. Because there were clones selected that are identical to a human mRNA in part but obviously not identical in the other part.

[0167] Also, if the finally selected clones were found to be not full-length compared with the sequences of human mRNA and ESTs, these clones were discarded.

#### EXAMPLE 11

50 A method for selection of clones by the combination of the ATGpr1 score and the prediction of the signal sequence by the PSORT (a method for selection of secretory proteins and membrane proteins that are novel and full-length).

[0168] The sequences of clones and the representative sequences of their clusters were used to obtain the maximal ATGpr1 score and predict the presence of the signal sequence. First, clones were selected based on the representative sequences of the clusters. The correspondence between the name and SEQ ID of the representative sequences used for selection (Table 368), and the correspondence between the name and SEQ ID of the introns (including the representative sequences of the 5'-end and 3'-end, and ESTs) used for selection of clones from the representative sequences of the groups (Table 369) were shown in the last part of the present specification. Therein, HRIFA and HRIRA indicate the representative sequence of the 5'-end group, and that of the 3'-end group, respectively.

[0169] In the clusters in which a single clone is contained (the sequence of the 5'-end clone = the representative sequence of the 5'-end), selected were the clones that were judged to be full-length in comparison with human mRNA and ESTs, having the maximal ATGpr1 score 0.5 or higher, and predicted to contain the signal sequence, in principle. However, in the following cases, a clone having a longer 5'-end was selected: the maximal ATGpr1 score was less than 0.5, the sequence of the 5'-end was not full-length, the clone was obviously shorter although the clone was not classified into the same cluster according to the BLAST search of the other clones, or the 5'-end sequence corresponding to the 3'-end of the other clones in the same cluster in which the 3'-end sequence of the clone was contained was found to be longer by assembling. Furthermore, if there were multiple full-length clones in the same cluster and it was not successful to determine by assembling which has the longer 5'-end, all the clones were selected. For assembling, the Sequencher<sup>™</sup> (Hitachi Soft Engineering) was used. As a result, the signal sequence predicted to be present in the representative sequence was not present in some of the selected clones. In some cases, the ATGpr1 score became smaller than 0.5 or 0.3. The fullness ratio in these clones was low, yet still it is possible that the clones are full-length. The clones in which the signal sequence predicted to be present in the representative sequence was not present after selection were likely to be without the signal sequence, but still it is possible that the clones encode a membrane protein.

[0170] In the clusters comprising multiple clones, in which the representative sequence of the 5'-end was predicted to contain the signal sequence, selected were the clones having the longest 5'-end sequence among the clones which were judged to be full-length compared with human mRNA and ESTs, having the maximal ATGpr1 score for the 5'-end sequence 0.5 or higher, and predicted to contain the signal sequence. However, in the following cases, a clone having a longer 5'-end was selected: the maximal ATGpr1 score was less than 0.5, the sequence of the 5'-end was not full-length, the clone was obviously shorter although the clone was not classified into the same cluster according to the BLAST search of the other clones, or the 5'-end sequence corresponding to the 3'-end of the other clones in the same cluster in which the 3'-end sequence of the clone was contained was found to be longer. Furthermore, if there were multiple full-length clones in the same cluster and it was not successful to determine by assembling which has the longer 5'-end, all the clones were selected. As a result, the signal sequence predicted to be present in the representative sequence was not present in some of the selected clones. In some cases, the ATGpr1 score became smaller than 0.5 or 0.3. The fullness ratio in these clones was low, yet still it is possible that the clones are full-length. The clones in which the signal sequence predicted to be present in the representative sequence was not present after selection were likely to be without the signal sequence at the 5'-end, but still it is possible that the clones encode a membrane protein.

[0171] Next, in the clusters comprising multiple clones, in which the representative sequence of the 5'-end was predicted to have no signal sequence, selected were the clones which were judged to be full-length compared with human mRNA and ESTs, having the maximal ATGpr1 score for the 5'-end sequence 0.5 or higher, and predicted to contain the signal sequence.

[0172] The number of the clones selected by the combination of the ATGpr1 score and the prediction of a signal sequence by the PSORT were 254. The number of the clones having the maximal ATGpr1 score 0.5 or higher, and predicted to contain a signal sequence were 170 (Table 7-10). Among the clones, 164 clones were found to have the representative sequence of the original cluster that fulfills the same conditions. On the other hand, 5 clones were selected from the representative sequences of the 5'-end of the clusters which was predicted to contain a signal sequence while the maximal ATGpr1 score was lower than 0.5. A clone was selected from the representative sequence of the 5'-end of the cluster which was predicted to have no signal sequence.

[0173] The clones that have the maximal ATGpr1 score 0.3 or higher and less than 0.5 and predicted to contain the signal sequence were 35 clones (Table 11), in which 8 clones were found to have the representative sequence of the original cluster that fulfills the same conditions. Twenty-seven clones were selected from the representative sequences of the clusters which have the maximal ATGpr1 score 0.5 or higher and were predicted to have no signal sequence.

[0174] The clones that have the maximal ATGpr1 score less than 0.3 and were predicted to contain a signal sequence were 41 clones (Table 12). The clones that have the maximal ATGpr1 score 0.5 or higher and were predicted to have no signal sequence were 4 clones (Table 13). The clones that have the maximal ATGpr1 score 0.3 or higher and less than 0.5 and were predicted to have no signal sequence were 2 clones (Table 14). The clones that have the maximal ATGpr1 score less than 0.3 and were predicted to contain a signal sequence were 2 clones (Table 15). The representative sequences of the original clusters of all the clones had the maximal ATGpr1 score 0.3 or higher, and were predicted to contain a signal sequence.

[0175] The fullness ratio of the clones having the maximal ATGpr1 score 0.5 or higher, 0.3 or higher, and 0 or higher is expected to be as shown in Table 3, 4, 5, and 6.

Table 7

Table 13

Four clones from which selected clones have the maximal ATGpr1 score 0.5 or higher, and predicted to be lacking the signal sequence by the PSORT

name of clone	name of sequence	maximal ATGpr1 score	signal	name of representative sequence	maximal ATGpr1 score	signal
NT2RP3002281	F-NT2RP3002281	0.81	No	HRIFA012999a	0.61	Yes
NT2RP3002721	F-NT2RP3002721	0.94	No	HRIFA023305a	0.57	Yes
NT2RP3004083	F-NT2RP3004083	0.94	No	HRIFA008387a	0.76	Yes
PLACE1005669	F-PLACE1005669	0.94	No	HRIFA012513a	0.65	Yes

Table 14

Two clones from which selected clones have the maximal ATGpr1 score 0.3 or higher and less than 0.5 and predicted to have no signal sequence by the PSORT

name of clone	name of sequence	maximal ATGpr1 score	signal	representative sequence	maximal ATGpr1 score	signal
NT2RP3000481	F-NT2RP3000481	0.47	No	HRIFA028614a	0.93	Yes
NT2RP3003559	F-NT2RP3003559	0.48	No	HRIFA025514a	0.45	Yes

Table 15

Two clones from which selected clones have the maximal ATGpr1 score 0 or higher and less than 0.3, and predicted to have no signal sequence by the PSORT

name of clone	name of sequence	maximal ATGpr1 score	signal	representative sequence	maximal ATGpr1 score	signal
PLACE1005601	F-PLACE1005601	0.12	No	HRIFA010593a	0.64	Yes
PLACE1006786	F-PLACE1006786	0.22	No	HRIFA012333a	0.51	Yes

#### EXAMPLE 12

A method for the selection of clones based on the top hit data in the homology search against the SwissProt (a method for the selection of a novel full-length protein that is predicted to have a function based on the top hit data).

[0176] The representative sequences of the clusters were discarded in which the 5'-end sequence is identical (90% or more matching), or not similar (the compared part contains a sequence of 25 nucleotides or shorter and the similarity is lower than 60%) to the top hit data in the SwissProt. Then, the remaining representative sequences which has similarity to the representative sequences of the 5'-ends were classified by a group of the above keywords (some representative sequences belong to a group by multiple keywords), and then clones were selected from the clusters. The names and the corresponding SEQ IDs of the representative sequences, and also the names of the introns (including the representative sequence of the 5'-end or the 3'-end, or ESTs) used for selecting the clones from the representative sequences and the corresponding SEQ IDs are shown in the last part of the present specification (Table 368 and 369, respectively). HRIFA indicates the representative sequence of the 5'-end group, and HRIRA indicates the representative sequence of the 3'-end group.

[0177] In principle, from the clusters containing only a single clone (the 5'-end sequence is the representative sequence of the cluster), the clone was selected. However, in the following cases, the clone containing a longer 5'-end was selected: where the maximal ATGpr1 score was less than 0.5, the 5'-end sequence of the clone to be selected was not complete, or the 5'-end of the clone was found to be obviously short nevertheless the clone should not be

included in the same cluster based on the BLAST analysis between the other clones, or further, the 5'-end sequence of the said clone, which corresponds to the 3'-ends of the other clones belonging to the same cluster in which the 3'-end of the said clone was included, was turn out to be longer than those of the other clones by assembling them. When there were two clones in the same cluster, judged to be full-length, and it was difficult to determine which clone has the longer 5'-end even by assembling them, all the clones were selected. As a result, the ATGpr1 score in some clones became less than 0.5 or less than 0.3. The fullness ratio of these clones became lower, but there is still a possibility that the clones are full-length.

**[0178]** In the case in which multiple clones were contained in a cluster, selected was the clone having the longest 5'-end in the clones judged to be full-length compared to the human mRNA or human EST. However, in the following cases, the clone containing a longer 5'-end was selected: where the maximal ATGpr1 score was less than 0.5, the 5'-end sequence of the clone to be selected was not complete, or the 5'-end of the clone was found to be obviously short nevertheless the clone should not be included in the same cluster based on the BLAST analysis between the other clones, or further, the 5'-end sequence of the said clone, which corresponds to the 3'-ends of the other clones belonging to the same cluster in which the 3'-end of the said clone was included, was turn out to be longer than those of the other clones by assembling them. When there were two clones in the same cluster, judged to be full-length, and it was difficult to determine which clone has the longer 5'-end even by assembling them, all the clones were selected. As a result, the ATGpr1 score in some clones became less than 0.5 or less than 0.3. These clones can still be full-length.

**[0179]** Based on the top hit data in the SwissProt homology search, 658 clones were selected. Among them, 446 clones were selected by the keywords, secretion or membrane. Using the keyword, glycoprotein, 243 clones were selected. 51 clones were selected by the keywords for signal transduction. With the keywords for transcription, 130 clones were selected. 17 clones were selected by the keywords for disease.

**[0180]** Among the 446 clones selected by the keywords, secretion or membrane, 77 clones were overlapped with those selected by combining the ATGpr1 score and prediction by the PSORT for the existence of a signal sequence. Also, many clones were overlapped with those selected by the keyword, glycoprotein. Moreover, some clones were overlapped with the clones selected by the keywords for diseases.

**[0181]** Among the 243 clones selected by the keyword, glycoprotein, 53 clones were overlapped with those selected by combining the ATGpr1 score and prediction by the PSORT for the existence of a signal sequence. Also, many clones were overlapped with those selected by the keywords, secretion or membrane. Moreover, some clones were overlapped with the clones selected by the keywords in diseases.

**[0182]** Among the clones selected by the top hit data in the homology search on the SwissProt, 532 clones were having the maximal ATGpr1 score 0.5 or higher. 59 clones were having the maximal score 0.3 or higher and less than 0.5. 67 clones were with the maximal score less than 0.3.

**[0183]** When the maximal ATGpr1 score is 0.5 or higher, 0.3 or higher, no less than 0, the expected fullness ratio is as shown in Table 3, 4, 5, and 6, respectively.

Table 16

The representative sequences of the most homologous sequences in the SwissProt with the keyword(s) "growth factor", "cytokine", or "hormone", and the selected clones.

name of clone	name of representative sequence
HEMBA1001563	HRIFA001439a
HEMBA1003047	HRIFA002743a
HEMBA1005070	HRIFA020144a
HEMBA1006724	HRIFA021620a
HEMBA1006916	HRIFA021855a
MAMMA1001066	HRIFA027355a
MAMMA1001634	HRIFA027187a
MAMMA1002165	HRIFA027673a
NT2RM4000326	HRIFA032530a
NT2RM4001377	HRIFA005300a
NT2RP2000447	HRIFA006448a
NT2RP2000663	HRIFA006609a
NT2RP2000903	HRIFA006798a
NT2RP2002974	HRIFA027860a
NT2RP2003369	HRIFA008596a
NT2RP2004141	HRIFA009123a

Table 19 (continued)

The representative sequences of the most homologous sequences in the SwissProt with the keyword(s) "transmembrane", and the selected clones (except overlapped with Table 16, 17, and 18)		
	name of clone	name of representative sequence
5	THYRO1000400	HRIFA000564a
	THYRO1000584	HRIFA029209a
	THYRO1000678	HRIFA029256a
10	THYRO1000776	HRIFA029317a
	THYRO1000795	HRIFA029327a
	THYRO1000866	HRIFA027714a
	THYRO1001113	HRIFA029460a
	THYRO1001128	HRIFA029467a
15	THYRO1001242	HRIFA032360a
	THYRO1001266	HRIFA030264a
	THYRO1001456	HRIFA030370a
	THYRO1001529	HRIFA030411a
20	THYRO1001702	HRIFA030511a
	Y79AA1000127	HRIFA026121a
	Y79AA1000270	HRIFA005644a
	Y79AA1001426	HRIFA028651a
	Y79AA1001787	HRIFA028790a
25	Y79AA1001799	HRIFA032070a
	Y79AA1002213	HRIFA032224a

Table 20

The representative sequences of the most homologous sequences in the SwissProt with the keyword(s) "signal", and the selected clones (except overlapped with Table 16, 17, 18 and 19)		
	name of clone	name of representative sequence
30		
35	BNGH41000087	HRIFA029508a
	HEMBA1000128	HRIFA000123a
	HEMBA1000443	HRIFA000415a
	HEMBA1000590	HRIFA000553a
40	HEMBA1000634	HRIFA004780a
	HEMBA1000745	HRIFA000695a
	HEMBA1001221	HRIFA001132a
	HEMBA1001228	HRIFA001138a
	HEMBA1001390	HRIFA000071a
45	HEMBA1002131	HRIFA001942a
	HEMBA1002167	HRIFA001975a
	HEMBA1002178	HRIFA001984a
	HEMBA1002524	HRIFA002284a
	HEMBA1002992	HRIFA002694a
50	HEMBA1003072	HRIFA002762a
	HEMBA1003315	HRIFA000016a
	HEMBA1003487	HRIFA003055a
	HEMBA1003530	HRIFA003093a
55	HEMBA1005145	HRIFA003946a
	HEMBA1005337	HRIFA019651a
	HEMBA1005449	HRIFA020707a
	HEMBA1005522	HRIFA021398a

Table 20 (continued)

The representative sequences of the most homologous sequences in the SwissProt with the keyword(s) "signal", and the selected clones (except overlapped with Table 16, 17, 18 and 19)		
	name of clone	name of representative sequence
5	HEMBA1006335	HRIFA012069a
	HEMBA1006572	HRIFA021543a
	HEMBA1006707	HRIFA021499a
10	HEMBA1006749	HRIFA021637a
	HEMBA1006902	HRIFA021754a
	HEMBA1007013	HRIFA021906a
	HEMBA1007057	HRIFA022985a
	HEMBB1000447	HRIFA001558a
15	HEMBB1000567	HRIFA029730a
	HEMBB1000881	HRIFA029932a
	HEMBB1001026	HRIFA030025a
	HEMBB1001048	HRIFA030045a
20	HEMBB1001847	HRIFA031249a
	MAMMA1000106	HRIFA024482a
	MAMMA1000226	HRIFA025978a
	MAMMA1000591	HRIFA026303a
	MAMMA1001043	HRIFA026764a
25	MAMMA1001957	HRIFA027536a
	MAMMA1002080	HRIFA016963a
	MAMMA1002234	HRIFA027722a
	MAMMA1002633	HRIFA030461a
30	MAMMA1003126	HRIFA029263a
	NT2RM1000462	HRIFA004426a
	NT2RM1000542	HRIFA004490a
	NT2RM2000410	HRIFA022055a
	NT2RM2000423	HRIFA022065a
35	NT2RM2000622	HRIFA022156a
	NT2RM2000773	HRIFA023894a
	NT2RM2001626	HRIFA028911a
	NT2RM2001818	HRIFA031935a
40	NT2RM4000648	HRIFA032730a
	NT2RM4001843	HRIFA024718a
	NT2RP1000050	HRIFA005102a
	NT2RP1001004	HRIFA005720a
	NT2RP2000394	HRIFA003640a
45	NT2RP2000514	HRIFA006494a
	NT2RP2001480	HRIFA007219a
	NT2RP2001755	HRIFA007424a
	NT2RP2001878	HRIFA007512a
	NT2RP2002188	HRIFA007745a
50	NT2RP2002564	HRIFA007244a
	NT2RP2002824	HRIFA008200a
	NT2RP2003042	HRIFA008362a
	NT2RP2003593	HRIFA008252a
55	NT2RP2003931	HRIFA008976a
	NT2RP2004606	HRIFA009451a
	NT2RP2004648	HRIFA009482a

Table 20 (continued)

The representative sequences of the most homologous sequences in the SwissProt with the keyword(s) "signal", and the selected clones (except overlapped with Table 16, 17, 18 and 19)

	name of clone	name of representative sequence
5	NT2RP2005163	HRIFA009825a
	NT2RP2005247	HRIFA009881a
	NT2RP2005378	HRIFA004919a
10	NT2RP2005541	HRIFA010090a
	NT2RP2005883	HRIFA010319a
	NT2RP2006042	HRIFA010425a
	NT2RP3000063	HRIFA022528a
	NT2RP3000436	HRIFA022776a
15	NT2RP3000444	HRIFA022782a
	NT2RP3000481	HRIFA028614a
	NT2RP3000721	HRIFA009825a
	NT2RP3001012	HRIFA023129a
20	NT2RP3001159	HRIFA023212a
	NT2RP3001592	HRIFA023464a
	NT2RP3001754	HRIFA007728a
	NT2RP3002311	HRIFA024718a
	NT2RP3002738	HRIFA020748a
25	NT2RP3002790	HRIFA026519a
	NT2RP3002887	HRIFA029278a
	NT2RP3003354	HRIFA008212a
	NT2RP3003448	HRIFA025479a
30	NT2RP3003473	HRIFA001413a
	NT2RP3003614	HRIFA032642a
	NT2RP3004075	HRIFA010301a
	NT2RP3004090	HRIFA027329a
	NT2RP3004202	HRIFA025327a
35	NT2RP3004309	HRIFA025778a
	NT2RP3004345	HRIFA025353a
	NT2RP3004557	HRIFA025907a
	NT2RP4001467	HRIFA013276a
40	OVARC1000313	HRIFA011016a
	OVARC1000410	HRIFA022691a
	OVARC1000439	HRIFA011105a
	OVARC1001086	HRIFA011580a
	OVARC1001569	HRIFA022728a
45	OVARC1001570	HRIFA019412a
	PLACE1001407	HRIFA012069a
	PLACE1001464	HRIFA013276a
	PLACE1001516	HRIFA012702a
50	PLACE1001795	HRIFA012885a
	PLACE1001918	HRIFA012969a
	PLACE1002080	HRIFA0130921a
	PLACE1002153	HRIFA013135a
	PLACE1002355	HRIFA013254a
55	PLACE1002374	HRIFA013265a
	PLACE1002726	HRIFA018688a
	PLACE1003428	HRIFA013911a

Table 20 (continued)

The representative sequences of the most homologous sequences in the SwissProt with the keyword(s) "signal", and the selected clones (except overlapped with Table 16, 17, 18 and 19)		
	name of clone	name of representative sequence
5	PLACE1003460	HRIFA013932a
	PLACE1003772	HRIFA014133a
	PLACE1004078	HRIFA014336a
10	PLACE1004520	HRIFA014621a
	PLACE1004648	HRIFA014702a
	PLACE1004887	HRIFA014868a
	PLACE1005426	HRIFA015246a
	PLACE1006071	HRIFA016639a
15	PLACE1006443	HRIFA015902a
	PLACE1006716	HRIFA016070a
	PLACE1006959	HRIFA016214a
	PLACE1007077	HRIFA016639a
20	PLACE1007081	HRIFA016290a
	PLACE1007702	HRIFA016669a
	PLACE1008657	HRIFA017257a
	PLACE1008744	HRIFA017312a
	PLACE1009546	HRIFA017801a
25	PLACE1011116	HRIFA018754a
	PLACE1011708	HRIFA019105a
	PLACE2000118	HRIFA024994a
	PLACE3000213	HRIFA015486a
30	PLACE4000354	HRIFA015486a
	THYRO1000061	HRIFA013279a
	THYRO1000846	HRIFA029349a
	THYRO1001063	HRIFA029434a
	THYRO1001608	HRIFA050456a
35	THYRO1001803	HRIFA030566a
	Y79AA1000876	HRIFA030629a
	Y79AA1001090	HRIFA028511a
	Y79AA1001272	HRIFA028576a
40	Y79AA1001727	HRIFA006642a
	Y79AA1001803	HRIFA032073a
	Y79AA1002376	HRIFA032820a

Table 21

The representative sequences of the most homologous sequences in the SwissProt with the keyword(s) "glycoprotein", and the selected clones		
	name of clone	name of representative sequence
50	BNGH41000087	HRIFA029508a
	BNGH41000091	HRIFA029511a
	HEMBA1000275	HRIFA000264a
	HEMBA1000349	HRIFA000327a
55	HEMBA1000590	HRIFA000553a
	HEMBA1000634	HRIFA004780a
	HEMBA1000835	HRIFA000776a
	HEMBA1000907	HRIFA000845a



Table 21 (continued)

The representative sequences of the most homologous sequences in the SwissProt with the keyword(s) "glycoprotein", and the selected clones		
	name of clone	name of representative sequence
5	HEMBA1001221	HRIFA001132a
	HEMBA1001228	HRIFA001138a
	HEMBA1001621	HRIFA001489a
10	HEMBA1002131	HRIFA001942a
	HEMBA1002164	HRIFA001972a
	HEMBA1002167	HRIFA001975a
	HEMBA1002178	HRIFA001984a
	HEMBA1002316	HRIFA002102a
15	HEMBA1002421	HRIFA005392a
	HEMBA1002767	HRIFA002503a
	HEMBA1003047	HRIFA002743a
	HEMBA1003101	HRIFA002787a
20	HEMBA1003230	HRIFA002891a
	HEMBA1003392	HRIFA002980a
	HEMBA1004250	HRIFA003504a
	HEMBA1004391	HRIFA020693a
	HEMBA1004444	HRIFA029285a
25	HEMBA1004454	HRIFA003592a
	HEMBA1004505	HRIFA003635a
	HEMBA1005449	HRIFA020707a
	HEMBA1005489	HRIFA024543a
30	HEMBA1005522	HRIFA021398a
	HEMBA1005545	HRIFA020272a
	HEMBA1006335	HRIFA012069a
	HEMBA1006572	HRIFA021543a
	HEMBA1006707	HRIFA021499a
35	HEMBA1006724	HRIFA021620a
	HEMBA1006749	HRIFA021637a
	HEMBA1006902	HRIFA021754a
	HEMBA1007057	HRIFA022985a
40	HEMBA1007332	HRIFA022493a
	HEMBB1000447	HRIFA001558a
	HEMBB1000567	HRIFA029730a
	HEMBB1000679	HRIFA029802a
	HEMBB1000881	HRIFA029932a
45	HEMBB1001048	HRIFA030045a
	HEMBB1002427	HRIFA005760a
	MAMMA1000106	HRIFA024482a
	MAMMA1000403	HRIFA026465a
50	MAMMA1000591	HRIFA026303a
	MAMMA1000681	HRIFA026364a
	MAMMA1000706	HRIFA026382a
	MAMMA1001043	HRIFA026764a
	MAMMA1001237	HRIFA026860a
55	MAMMA1001615	HRIFA022865a
	MAMMA1001893	HRIFA027485a
	MAMMA1001978	HRIFA027549a

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Table 25 (continued)

The clones selected by the keyword(s) of the top hit data in the SwissProt, and having the maximal score in the ATGpr1 0.5 or higher.			
	name of clone	name of sequence	maximal ATGpr1 score
5	HEMBA1005522	F-HEMBA1005522	0.73
	HEMBA1005545	F-HEMBA1005545	0.94
	HEMBA1005698	F-HEMBA1005698	0.68
10	HEMBA1005929	F-HEMBA1005929	0.72
	HEMBA1005945	F-HEMBA1005945	0.80
	HEMBA1006276	F-HEMBA1006276	0.50
	HEMBA1006299	F-HEMBA1006299	0.94
	HEMBA1006335	F-HEMBA1006335	0.94
15	HEMBA1006430	F-HEMBA1006430	0.93
	HEMBA1006482	F-HEMBA1006482	0.59
	HEMBA1006517	F-HEMBA1006517	0.68
	HEMBA1006544	F-HEMBA1006544	0.94
20	HEMBA1006572	F-HEMBA1006572	0.62
	HEMBA1006707	F-HEMBA1006707	0.94
	HEMBA1006724	F-HEMBA1006724	0.80
	HEMBA1006749	F-HEMBA1006749	0.94
	HEMBA1006902	F-HEMBA1006902	0.94
25	HEMBA1006916	F-HEMBA1006916	0.80
	HEMBA1007013	F-HEMBA1007013	0.82
	HEMBA1007057	F-HEMBA1007057	0.94
	HEMBA1007226	F-HEMBA1007226	0.50
30	HEMBA1007241	F-HEMBA1007241	0.94
	HEMBA1000106	F-HEMBA1000106	0.94
	HEMBA1000447	F-HEMBA1000447	0.73
	HEMBA1000668	F-HEMBA1000668	0.50
	HEMBA1000679	F-HEMBA1000679	0.91
35	HEMBA1000881	F-HEMBA1000881	0.77
	HEMBA1001026	F-HEMBA1001026	0.94
	HEMBA1001048	F-HEMBA1001048	0.88
	HEMBA1001200	F-HEMBA1001200	0.81
40	HEMBA1001573	F-HEMBA1001573	0.80
	HEMBA1001847	F-HEMBA1001847	0.81
	HEMBA1001959	F-HEMBA1001959	0.94
	HEMBA1002041	F-HEMBA1002041	0.79
	HEMBA1002051	F-HEMBA1002051	0.60
45	HEMBA1002302	F-HEMBA1002302	0.89
	HEMBA1002427	F-HEMBA1002427	0.94
	HEMBA1002661	F-HEMBA1002661	0.94
	MAMMA1000106	F-MAMMA1000106	0.78
50	MAMMA1000204	F-MAMMA1000204	0.94
	MAMMA1000226	F-MAMMA1000226	0.94
	MAMMA1000403	F-MAMMA1000403	0.59
	MAMMA1000473	F-MAMMA1000473	0.86
	MAMMA1000496	F-MAMMA1000496	0.70
55	MAMMA1000591	F-MAMMA1000591	0.77
	MAMMA1000681	F-MAMMA1000681	0.94
	MAMMA1000788	F-MAMMA1000788	0.83

**EXAMPLE 13**

## Selection of cDNA clone NT2RP2036580

- 5 **[0184]** Clone NT2RP2006580 as well as clone HEMBA1000121 was selected from the representative sequences belonging to HRIFA000116a cluster of the most homologous sequence in the SwissProt with the keywords "trans-membrane". Although each of the clones, HEMBA1000121 and NT2RP2006580, was assembled with other clones for 5' extension, any other clones did not extend the clones toward the 5' direction. Accordingly, it is possible that both clones are full-length cDNA clones. The maximal ATGpr1 score of F-NT2RP2006580 is 0.37, and therefore, the fullness ratio is low. However, it is still possible for the sequence to cover the full-length.
- 10 **[0185]** Thus, the total number of selected clones is 830. Based on the top matching data resulted from Swiss-Prot homology search, 659 clones were selected. From them, 447 clones were selected by the keywords of "secretion" and "membrane". Among the clones selected based on the top matching data, 60 clones exhibited the maximal ATGpr1 score of 0.3 or higher and less than 0.5.
- 15 **[0186]** The sequences of F-NT2RP2006580 and R-NT2RP2006580 are shown in SEQ ID NO: 2545 and SEQ ID NO: 2546, respectively.

**EXAMPLE 14**

## 20 Full-length sequence analysis and homology search

- [0187]** Full-length sequence was determined for each selected cDNA clones. The nucleotide sequence determination was performed mainly by the dye-terminator method using custom synthesized DNA primers according to the primer walking procedure (custom synthesized DNA primers were used for sequencing; sequencing reaction was performed with DNA sequencing reagent supplied by PE Biosystems according to the supplier's manual; and the samples were analyzed in an automatic sequencer made by the same supplier). Sequence determination of some clones was carried out in the same manner but using a Licor DNA sequencer. Overlapping partial nucleotide sequences, which were obtained by the above-described method, were assembled together to determine a full-length nucleotide sequence. Amino acid sequences were then deduced from the determined full-length nucleotide sequences. However, amino acid sequence is not shown for a clone of which coding region was hard to be deduced or of which amino acid sequence has less than 100 amino acid residues. SEQ ID NOs corresponding to the respective clones are indicated in Table 370.
- 25 **[0188]** GenBank, Swiss-Prot and UniGene were searched for the determined nucleotide sequences by BLAST analysis. Matching data of cDNA clone which exhibits higher homology and of which functions are easily predicted based on the nucleotide sequences and the deduced amino acid sequences are selected from the BLAST analysis matching data with P value of  $10^{-4}$  or less. The matching data selected are listed herein. However, there are some clones that did not match the criteria for judgment and such matching data of BLAST analysis are not shown herein. The results of homology search indicated in the last part of this specification are as follows.
- 30 **[0189]** Homology search result 1: data obtained by the homology search of Swiss-Prot database for representative sequences of the 5'-end cluster
- [0190]** Homology search result 2: homology of representative sequences of the 5'-end cluster to the data in Swiss-Prot database; the P value is  $10^{-10}$  or less
- [0191]** Homology search result 3: homology of representative sequences of the 5'-end cluster to the data in Swiss-Prot database; the P value is higher than  $10^{-10}$  and  $10^{-4}$  or less
- 35 **[0192]** Homology search result 4: homology of representative sequences of the 5'-end cluster to the data in Swiss-Prot database; the P value is higher than  $10^{-4}$  and 1 or less
- [0193]** Homology search result 5: data obtained by the homology search of Swiss-Prot database for 5'-end sequences of cDNA clone
- [0194]** Homology search result 6: data obtained by the homology search of GenBank database (<http://www.ncbi.nlm.nih.gov/web/GenBank/>) except for EST and STS sequence data for 5'-end sequences of cDNA clone
- 40 **[0195]** Homology search result 7: data obtained by the homology search of GenBank database (<http://www.ncbi.nlm.nih.gov/web/GenBank/>) except for EST and STS sequence data for 3'-end sequences of cDNA clone
- [0196]** Homology search result 8: data obtained by the homology search of Human UniGene database (<http://www.ncbi.nlm.nih.gov/Unigene/>) for 5'-end sequences of cDNA clone
- [0197]** Homology search result 9: data obtained by the homology search of Human UniGene database (<http://www.ncbi.nlm.nih.gov/Unigene/>) for 3'-end sequences of cDNA clone
- 45 **[0198]** Homology search result 10: result obtained by the homology search for full-length nucleotide sequences and deduced amino acid sequences
- [0199]** The P value indicates similarity between two sequences as a score by considering the probability that the two

sequences are accidentally similar. In general, as the value is lower, the similarity is higher. In general, as the value is lower, the homology is higher (Altschul, S.F., Gish, W., Miller, W., Myers, E.W. & Lipman, D.L. (1990) "Basic local alignment search tool." *J. Mol. Biol.* 215:403-410; Gish, W. & States, D.J. (1993) "Identification of protein coding regions by database similarity search." *Nature Genet.* 3:266-272).

#### Example 15. Gene expression analysis with hybridization using high density DNA filter

**[0200]** Nylon membrane for DNA spotting was prepared according to the following procedure. *E. coli* was cultured in each well of a 96-well plate (in a LB medium at 37. for 16 hours). A sample of each culture was suspended in 10 . 1 of sterile water in a well of a 96-well plate. The plate was heated at 100. for 10 minutes. Then, the boiled samples were analyzed by PCR. PCR was performed in a 20 .1 solution by using TaKaRa PCR Amplification Kit (Takara) according to the supplier's protocol. Primers used for the amplification of an insert cDNA in a plasmid were a pair of sequencing primers, ME761FW (5' tacggaagtgttactctgc 3' / SEQ ID NO: 3591) and ME1250RV (5' tgtgggagggttttctcta 3' / SEQ ID NO: 3592), or a pair of primers, M13M4 (5' gtttcccgatcagcagc 3' / SEQ ID NO: 3593) and M13RV (5' cag-gaaacagctatgac 3' / SEQ ID NO: 3594). PCR was performed using a thermal cycler, GeneAmp System 9600 (PE Biosystems) at 95. for 5 minutes; at 95. for 10 seconds and at 68. for 1 minute for 10 cycles; at 98. for 20 seconds and at 60. for 3 minutes for 20 cycles; and at 72. for 10 minutes. After the PCR, the 20 .1 reaction solution was loaded onto a 1% agarose gel and fractionated by electrophoresis. DNA on the gel was stained with ethidium bromide to confirm the amplification of cDNA. When cDNAs were not amplified by PCR, plasmids containing the corresponding insert cDNAs were prepared by the alkali-extraction method (J. Sambrook, E.F., Fritsh, & T. Maniatis, "Molecular Cloning, A laboratory manual/ 2nd edition, Cold Spring Harbor Laboratory Press, 1989).

**[0201]** Preparation of DNA array was carried out by the following procedure. A sample of a DNA solution was added in each well of a 384-well plate. DNA was spotted onto a nylon membrane (Boehringer) by using a 384-pin tool of Biomek 2000 Laboratory Automation System (Beckman-Coulter). Specifically, the 384-well plate containing the DNA was placed under the 384-pin tool. The independent 384 needles were simultaneously dipped into the DNA solution for DNA deposition. The needles were gently pressed onto a nylon membrane and the DNA deposited at the tips of needles was spotted onto the membrane. Denaturation of the spotted DNA and immobilization of the DNA on the nylon membrane were carried out according to standard methods (J. Sambrook, E.F., Fritsh, & T. Maniatis, "Molecular Cloning, A laboratory manual/ 2nd edition, Cold Spring Harbor Laboratory Press, 1989).

**[0202]** A probe for hybridization was radioisotope-labeled first strand cDNA. Synthesis of the first strand cDNA was performed by using Thermoscript<sup>TM</sup> RT-PCR System (GIBCO). Specifically, the first strand cDNA was synthesized by using 1.5 .g of mRNAs from various human tissues (Clontech), 1 .1 of 50.M Oligo(dT)20 and 50.Ci [<sup>32</sup>P]dATP according to an attached protocol. Purification of a probe was carried out by using ProbeQuant<sup>TM</sup> G-50 micro column (Amersham-Pharmacia Biotech) according to an attached protocol. In the next step, 2 units of *E. coli* RNase H were added to the reaction mixture. The mixture was incubated at room temperature for 10 minutes, and then, 100.g of human COT-1 DNA (GIBCO) was added thereto. The mixture was incubated at 97. for 10 minutes and then was allowed to stand on ice to give hybridization probe.

**[0203]** Hybridization of the radioisotope-labeled probe to the DNA array was performed according to standard methods (J. Sambrook, E.F., Fritsh, & T. Maniatis, Molecular Cloning, A laboratory manual/ 2nd edition, Cold Spring Harbor Laboratory Press, 1989). The membrane was washed as follows: the nylon membrane was washed 3 times by incubating it in Washing solution 1 (2xSSC, 1% SDS) at room temperature (about 26.) for 20 minutes; then the membrane was washed 3 times by incubating it in Washing solution 2 (0.1xSSC, 1% SDS) at 65. for 20 minutes.

**[0204]** Autoradiography was performed by using an image plate for BAS2000 (Fuji Photo Film Co., Ltd.). Specifically, the nylon membrane with probe hybridized thereon was wrapped with a piece of Saran Wrap and brought into tight contact with the image plate on the light-sensitive surface. The membrane with the image plate was placed in an imaging cassette for radioisotope and allowed to stand in dark place for 4 hours. The radioactivity recorded on the image plate was analyzed by using BAS2000 (Fuji Photo Film Co., Ltd.). The activity was subjected to electronic conversion and recorded as an image file of autoradiogram. The signal intensity of each DNA spot was analyzed by using Visage High Density Grid Analysis Systems (Genomic Solutions Inc.). The signal intensity was converted into numerical data. The data were taken in duplicate. The reproducibility was assessed by comparing the signal intensities of the corresponding spots on the duplicated DNA filters that were hybridized to a single DNA probe (Figure 2). In 95% of entire spots, the ratio between the corresponding spots falls within a range of 2 or less, and the correlation coefficient is  $r=1.97$ . Thus, the reproducibility is satisfactory.

**[0205]** The detection sensitivity in gene expression analysis was estimated by examining increases in the signal intensity of probe concentration-dependent spot in hybridization using a probe complementary to the DNA spotted on the nylon membrane. DNA used was PLACE 1008092 (the same as DNA deposited in GenBank under an Accession No. AF107253). The DNA array with DNA of PLACE1008092 was prepared according to the above-mentioned method. The probe used was prepared as follows: mRNA was synthesized in vitro from the clone, PLACE1008092. By using

this mRNA as a template, radioisotope-labeled first strand cDNA was synthesized in the same manner as described above, and the cDNA was used as the probe. In order to synthesize mRNA from PLACE1008092 in vitro, a plasmid in which the 5' end of the cDNA PLACE1008092 was ligated to the T7 promoter of pBluescript SK(-) was constructed. Specifically, the PLACE1008092 insert was cut out from pME18SFL3 carrying the cDNA at a DraIII site thereof by XhoI digestion. The resulting PLACE1008092 fragment was ligated to XhoI-predigested pBluescript SK(-) by using DNA ligation kit ver.2 (Takara). The in vitro mRNA synthesis from PLACE1008092 inserted into pBluescript SK(-) was carried out by using Ampliscribe<sup>TM</sup> T7 high yield transcription kit (Epicentre technologies). Hybridization and the analysis of signal intensity of each DNA spot were performed by the same methods as described above. When the probe concentration is  $1 \times 10^7$  g/ml or less, there was no increase of signal intensity proportional to the probe concentration. Therefore, it was assumed to be difficult to compare the signals with one another in this concentration range. Thus, the spots with the intensity of 40 or less were uniformly taken as low level signals (Figure 3). Within a concentration of the probe ranging from  $1 \times 10^7$  g/ml to 0.1 g/ml, the signal was found to increase in a probe concentration-dependent manner. The detection limit represented as the ratio of the expression level of test mRNA to that of total mRNA in a sample was 1:100,000.

[0206] Tables 28-184 (also containing clones without description in Examples) show the expression of each cDNA in human normal tissues (heart, lung, pituitary gland, thymus, brain, kidney, liver and spleen). The expression levels are indicated with numerical values of 0-10,000. Genes that were expressed in at least a single tissue are indicated below by the corresponding clone names:

	clone:	BNGH41000020,	BNGH41000087,	BNGH41000091,	HEMBA1000121,	HEMBA1000275,
20	HEMBA1000300,	HEMBA1000443,	HEMBA1000462,	HEMBA1000477,	HEMBA1000634,	HEMBA1000713,
	HEMBA1000835,	HEMBA1000875,	HEMBA1000940,	HEMBA1000962,	HEMBA1001228,	HEMBA1001296,
	HEMBA1001390,	HEMBA1001563,	HEMBA1001621,	HEMBA1002048,	HEMBA1002131,	HEMBA1002163,
	HEMBA1002164,	HEMBA1002167,	HEMBA1002178,	HEMBA1002195,	HEMBA1002227,	HEMBA1002239,
	HEMBA1002316,	HEMBA1002421,	HEMBA1002524,	HEMBA1002551,	HEMBA1002767,	HEMBA1002985,
25	HEMBA1002992,	HEMBA1003047,	HEMBA1003072,	HEMBA1003101,	HEMBA1003120,	HEMBA1003230,
	HEMBA1003294,	HEMBA1003315,	HEMBA1003392,	HEMBA1003399,	HEMBA1003487,	HEMBA1003530,
	HEMBA1003945,	HEMBA1004007,	HEMBA1004067,	HEMBA1001085,	HEMBA1004110,	HEMBA1004391,
	HEMBA1004444,	HEMBA1004454,	HEMBA1004505,	HEMBA1004797,	HEMBA1004952,	HEMBA1005070,
	HEMBA1005084,	HEMBA1005145,	HEMBA1005230,	HEMBA1005246,	HEMBA1005337,	HEMBA1005430,
30	HEMBA1005449,	HEMBA1005489,	HEMBA1005545,	HEMBA1005698,	HEMBA1005929,	HEMBA1005945,
	HEMBA1005016,	HEMBA1006171,	HEMBA1006276,	HEMBA1006311,	HEMBA1006335,	HEMBA1006357,
	HEMBA1006430,	HEMBA1006482,	HEMBA1006517,	HEMBA1006544,	HEMBA1006658,	HEMBA1006707,
	HEMBA1006749,	HEMBA1006770,	HEMBA1006902,	HEMBA1006912,	HEMBA1006916,	HEMBA1006960,
	HEMBA1007013,	HEMBA1007057,	HEMBA1007063,	HEMBA1007291,	HEMBA1007332,	HEMBA1007106,
35	HEMBA1000309,	HEMBA1000447,	HEMBA1000542,			
	HEMBA1000567,	HEMBA1000642,	HEMBA1000905,	HEMBA1001026,	HEMBA1001048,	HEMBA1001407,
	HEMBA1001530,	HEMBA1001573,	HEMBA1001847,	HEMBA1001959,	HEMBA1001978,	HEMBA1002039,
	HEMBA1002041,	HEMBA1002051,	HEMBA1002162,	HEMBA1002228,	HEMBA1002302,	HEMBA1002427,
	HEMBA1002465,	HEMBA1002661,	HEMBA1002663,	HEMBA1002693,	MAMMA1000046,	MAMMA1000102,
40	MAMMA1000106,	MAMMA1000118,	MAMMA1000204,	MAMMA1000226,	MAMMA1000403,	MAMMA1000449,
	MAMMA1000457,	MAMMA1000473,	MAMMA1000528,	MAMMA1000591,	MAMMA1000614,	MAMMA1000652,
	MAMMA1000681,	MAMMA1000706,	MAMMA1000788,	MAMMA1000810,	MAMMA1000814,	MAMMA1000881,
	MAMMA1000986,	MAMMA1000994,	MAMMA1001043,	MAMMA1001066,	MAMMA1001094,	MAMMA1001141,
	MAMMA1001150,	MAMMA1001284,	MAMMA1001310,	MAMMA1001344,	MAMMA1001418,	MAMMA1001532,
45	MAMMA1001609,	MAMMA1001615,	MAMMA1001634,	MAMMA1001893,	MAMMA1001901,	MAMMA1001957,
	MAMMA1002070,	MAMMA1002091,	MAMMA1002095,	MAMMA1002128,	MAMMA1002142,	MAMMA1002165,
	MAMMA1002205,	MAMMA1002224,	MAMMA1002586,	MAMMA1003126,	NT2RM1000407,	NT2RM1000462,
	NT2RM1000542,	NT2RM1000789,	NT2RM1000855,	NT2RM1000858,	NT2RM2000241,	NT2RM2000306,
	NT2RM2000410,	NT2RM2000423,	NT2RM2000497,	NT2RM2000514,	NT2RM2000565,	NT2RM2000582,
50	NT2RM2000589,	NT2RM2000622,	NT2RM2000773,	NT2RM2001126,	NT2RM2001626,	NT2RM2001792,
	NT2RM2001941,	NT2RM4000198,	NT2RM4000295,	NT2RM4000444,	NT2RM4000593,	NT2RM4000761,
	NT2RM4000965,	NT2RM4000997,	NT2RM4001321,	NT2RM4001325,		
	NT2RM4001377,	NT2RM4001735,	NT2RM4001768,	NT2RM4001843,	NT2RP1000002,	NT2RP1000181,
	NT2RP1000271,	NT2RP1000300,	NT2RP1000325,	NT2RP1000465,	NT2RP1000468,	NT2RP1000740,
55	NT2RP1000903,	NT2RP1000981,	NT2RP2000092,	NT2RP2000178,	NT2RP2000240,	NT2RP2000447,
	NT2RP2000479,	NT2RP2000533,	NT2RP2000610,	NT2RP2000616,	NT2RP2000694,	NT2RP2000739,
	NT2RP2001200,	NT2RP2001223,	NT2RP2001388,	NT2RP2001469,	NT2RP2001480,	NT2RP2001514,
	NT2RP2001529,	NT2RP2001538,	NT2RP2001562,	NT2RP2001662,	NT2RP2001878,	NT2RP2001903,

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	NT2RP2001921,	NT2RP2001956,	NT2RP2002015,	NT2RP2002063,	NT2RP2002188,	NT2RP2002232,
	NT2RP2002409,	NT2RP2002510,	NT2RP2002527,	NT2RP2002533,	NT2RP2002564,	NT2RP2002721,
	NT2RP2002824,	NT2RP2002942,	NT2RP2002974,	NT2RP2003138,	NT2RP2003179,	NT2RP2003210,
	NT2RP2003302,	NT2RP2003369,	NT2RP2003383,	NT2RP2003390,	NT2RP2003469,	NT2RP2003593,
5	NT2RP2003599,	NT2RP2003655,	NT2RP2003940,	NT2RP2003950,	NT2RP2004069,	NT2RP2004108,
	NT2RP2004141,	NT2RP2004179,	NT2RP2004205,	NT2RP2004447,	NT2RP2004524,	NT2RP2004556,
	NT2RP2004606,	NT2RP2004648,	NT2RP2004794,	NT2RP2004837,	NT2RP2004847,	NT2RP2005027,
	NT2RP2005069,	NT2RP2005163,	NT2RP2005181,	NT2RP2005247,	NT2RP2005378,	NT2RP2005391,
	NT2RP2005425,	NT2RP2005463,	NT2RP2005535,	NT2RP2005541,	NT2RP2005597,	NT2RP2005632,
10	NT2RP2005666,	NT2RP2005774,	NT2RP2005878,	NT2RP2005887,	NT2RP2005941,	NT2RP2006004,
	NT2RP2006042,	NT2RP2006092,	NT2RP2006099,	NT2RP2006269,	NT2RP3000011,	NT2RP3000022,
	NT2RP3000059,	NT2RP3000063,	NT2RP3000125,	NT2RP3000148,	NT2RP3000171,	NT2RP3000172,
	NT2RP3000201,	NT2RP3000232,	NT2RP3000304,	NT2RP3000378,	NT2RP3000436,	NT2RP3000460,
	NT2RP3000645,	NT2RP3000652,	NT2RP3000676,	NT2RP3000677,	NT2RP3000721,	NT2RP3000789,
15	NT2RP3000818,	NT2RP3000820,	NT2RP3000838,	NT2RP3000907,	NT2RP3000921,	NT2RP3001044,
	NT2RP3001159,	NT2RP3001170,	NT2RP3001195,	NT2RP3001271,	NT2RP3001388,	NT2RP3001560,
	NT2RP3001592,	NT2RP3001685,	NT2RP3001738,	NT2RP3001754,	NT2RP3001858,	NT2RP3001976,
	NT2RP3002015,	NT2RP3002160,	NT2RP3002281,	NT2RP3002311,	NT2RP3002324,	NT2RP3002353,
20	NT2RP3002409,	NT2RP3002411,	NT2RP3002721,	NT2RP3002737,	NT2RP3002738,	NT2RP3002836,
	NT2RP3002900,	NT2RP3002958,	NT2RP3003000,	NT2RP3003076,	NT2RP3003354,	NT2RP3003448,
	NT2RP3003469,	NT2RP3003473,	NT2RP3003532,	NT2RP3003614,	NT2RP3003729,	NT2RP3003849,
	NT2RP3003874,	NT2RP3003939,	NT2RP3003963,	NT2RP3004025,	NT2RP3004067,	NT2RP3004083,
	NT2RP3004090,	NT2RP3004119,	NT2RP3004130,	NT2RP3004133,	NT2RP3004202,	NT2RP3004294,
	NT2RP3004309,	NT2RP3004321,	NT2RP3004355,	NT2RP3004374,	NT2RP3004406,	NT2RP3004481,
25	NT2RP3004552,	NT2RP3004557,	NT2RP3004625,	NT2RP3004640,	NT2RP3004647,	NT2RP4000108,
	NT2RP4000634,	NT2RP4001877,	NT2RP4001879,	NT2RP4002187,	NT2RP4002715,	NT2RP4002750,
	OVARC1000090,	OVARC1000105,	OVARC1000137,	OVARC1000208,	OVARC1000255,	OVARC1000313,
	OVARC1000331,	OVARC1000410,	OVARC1000439,	OVARC1000467,	OVARC1000529,	OVARC1000553,
	OVARC1000775,	OVARC1000853,	OVARC1000873,	OVARC1000916,	OVARC1000956,	OVARC1000995,
30	OVARC1001030,	OVARC1001049,	OVARC1001086,	OVARC1001163,	OVARC1001260,	OVARC1001336,
	OVARC1001569,	OVARC1001570,	OVARC1001596,	OVARC1001807,	OVARC1001833,	OVARC1001991,
	PLACE1000231,	PLACE1000258,	PLACE1000442,	PLACE1000560,	PLACE1000912,	PLACE1000927,
	PLACE1001016,	PLACE1001100,	PLACE1001114,	PLACE1001183,	PLACE1001229,	PLACE1001340,
	PLACE1001407,	PLACE1001500,	PLACE1001516,	PLACE1001655,	PLACE1001836,	PLACE1001918,
35	PLACE1002080,	PLACE1002095,	PLACE1002153,	PLACE1002329,	PLACE1002374,	PLACE1002518,
	PLACE1002547,	PLACE1002726,	PLACE1002905,	PLACE1002911,	PLACE1002967,	PLACE1003163,
	PLACE1003407,	PLACE1003428,	PLACE1003438,	PLACE1003460,	PLACE1003529,	PLACE1003598,
	PLACE1003644,	PLACE1003772,	PLACE1003839,	PLACE1003845,	PLACE1003852,	PLACE1004078,
40	PLACE1004166,	PLACE1004168,	PLACE1004199,	PLACE1004279,	PLACE1004282,	PLACE1004305,
	PLACE1004441,	PLACE1004482,	PLACE1004492,	PLACE1004520,	PLACE1004630,	PLACE1004637,
	PLACE1004648,	PLACE1004816,	PLACE1004887,	PLACE1005005,	PLACE1005031,	PLACE1005383,
	PLACE1005410,	PLACE1005426,	PLACE1005539,	PLACE1005544,	PLACE1005569,	PLACE1005725,
	PLACE1005736,	PLACE1005768,	PLACE1005815,	PLACE1005878,	PLACE1005927,	PLACE1006071,
45	PLACE1006073,	PLACE1006079,	PLACE1006277,	PLACE1006443,	PLACE1006716,	PLACE1006809,
	PLACE1007077,	PLACE1007096,	PLACE1007626,	PLACE1007702,	PLACE1008469,	PLACE1008985,
	PLACE1009067,	PLACE1009527,	PLACE1009982,	PLACE1010078,	PLACE1010251,	PLACE1010445,
	PLACE1011045,	PLACE1011116,	PLACE1011181,	PLACE1011236,	PLACE1011364,	PLACE1011516,
	PLACE1011708,	PLACE1011978,	PLACE2000118,	PLACE2000219,	PLACE3000181,	PLACE4000354,
50	PLACE4000455,	SKNMC1000014,	THYRO1000061,	THYRO1000099,	THYRO1000584,	THYRO1000795,
	THYRO1000866,	THYRO1000999,	THYRO1001063,	THYRO1001113,	THYRO1001128,	THYRO1001205,
	THYRO1001237,	THYRO1001242,	THYRO1001456,	THYRO1001457,	THYRO1001478,	THYRO1001495,
	THYRO1001523,	THYRO1001529,	THYRO1001593,	THYRO1001608,	THYRO1001700,	THYRO1001702,
	THYRO1001725,	THYRO1001770,	THYRO1001803,	Y79AA1000127,	Y79AA1000207,	Y79AA1000226,
	Y79AA1000270,	Y79AA1000426,	Y79AA1000521,	Y79AA1000776,	Y79AA1000777,	Y79AA1000888,
55	Y79AA1000967,	Y79AA1001013,	Y79AA1001090,	Y79AA1001272,	Y79AA1001328,	Y79AA1001426,
	Y79AA1001427,	Y79AA1001430,	Y79AA1001523,	Y79AA1001530,	Y79AA1001592,	Y79AA1001727,
	Y79AA1001787,	Y79AA1001793,	Y79AA1001799,	Y79AA1001803,	Y79AA1001863,	Y79AA1002022,
	Y79AA1002213,	Y79AA1002373,	Y79AA1002376,	Y79AA1002381,		

**[0207]** Genes that were expressed in all the tissues tested are indicated below by the corresponding clone names: clone: BNGH41000020, HEMBA1000300, HEMBA1001390, HEMBA1002239, HEMBA1002316, HEMBA1004007, HEMBA1004067, HEMBA1005145, HEMBA1005230, HEMBA1005929, HEMBA1006357, HEMBA1006482, HEMBB1000567, HEMBB1001847, NEMBB1001978, MAMMA1000614, MAMMA1000652, MAMMA1000810, MAMMA1000814, MAMMA1001066, MAMMA1001094, MAMMA1001284, MAMMA1001310, MAMMA1001634, MAMMA1002165, MAMMA1002205, MAMMA1002224, NT2RM1000462, NT2RM1000855, NT2RM1000858, NT2RP2000423, NT2RM4000761, NT2RM4000997, NT2RP1000271, NT2RP1000325, NT2RP1000465, NT2RP2001538, NT2RP2001662, NT2RP2001903, NT2RP2002015, NT2RP2002188, NT2RP2002409, NT2RP2002510, NT2RP2002533, NT2RP2004556, NT2RP2004794, NT2RP2004847, NT2RP2005069, NT2RP2005163, NT2RP2005535, NT2RP2006269, NT2RP3000171, NT2RP3000645, NT2RP3000838, NT2RP3001271, NT2RP3001754, NT2RP3003076, NT2RP3003354, NT2RP3003614, NT2RP3004640, NT2RP3004647, OVARC1000090, OVARC1000208, OVARC1000553, OVARC1000995, OVARC1001030, OVARC1001049, PLACE1000231, PLACE1000258, PLACE1001516, PLACE1002080, PLACE1002911, PLACE1003598, PLACE1004648, PLACE1006443, PLACE1008469, PLACE1011708, PLACE2000118, THYRO1001128, THYRO1001205, THYRO1001242, THYRO1001803, Y79AA1000207, Y79AA1001013, Y79AA1001272, Y79AA1001328, Y79AA1001793, Y79AA1001863, Y79AA1002022, Y79AA1002376.

**[0208]** Genes that were expressed at low levels in any of the tissues tested are indicated below by the corresponding clone names: clone: HEMBA1000006, HEMBA1000128, HEMBA1000349, HEMBA1000590, HEMBA1000671, HEMBA1000732, HEMBA1000745, HEMBA1000907, HEMBA1001184, HEMBA1001221, HEMBA1001272, HEMBA1001297, HEMBA1001878, HEMBA1001886, HEMBA1002420, HEMBA1003497, HEMBA1003602, HEMBA1003732, HEMBA1004250, HEMBA1004785, HEMBA1004971, HEMBA1004982, HEMBA1005267, HEMBA1005522, HEMBA1005913, HEMBA1006299, HEMBA1006572, HEMBA1006724, HEMBA1007241, HEMBB1000276, HEMBB1000407, HEMBB1000668, HEMBB1000679, HEMBB1000881, HEMBB1001200, HEMBB1001547, HEMBB1002120, HEMBB1002245, MAMMA1000141, MAMMA1000496, MAMMA1001237, MAMMA1001623, MAMMA1001978, MAMMA1002080, MAMMA1002087, MAMMA1002234, MAMMA1002633, NT2RM1000580, NT2RM1000899, NT2RM2000632, NT2RM2001643, NT2PM2001818, NT2RM2001902, NT2RM2001939, NT2RM4000100, NT2RM4000115, NT2RM4000284, NT2RM4000326, NT2RM4000417, NT2RM4000587, NT2RM4000648, NT2RM4002352, NT2RP1000050, NT2RP1000239, NT2RP1000261, NT2RP1000448, NT2RP1000551, NT2RP1000579, NT2RP1000613, NT2RP1000679, NT2RP1001004, NT2RP1001020, NT2RP1001031, NT2RP1001563, NT2RP2000394, NT2RP2000514, NT2RP2000649, NT2RP2000663, NT2RP2000712, NT2RP2000818, NT2RP2000903, NT2RP2001276, NT2RP2001495, NT2RP2001755, NT2RP2001769, NT2RP2001817, NT2RP2001915, NT2RP2001948, NT2RP2002304, NT2RP2002674, NT2RP2002976, NT2RP2003042, NT2RP2003545, NT2RP2003664, NT2RP2003931, NT2RP2004495, NT2RP2004670, NT2RP2005514, NT2RP2005883, NT2RP2005994, NT2RP2006134, NT2RP2006512, NT2RP3000169, NT2RP3000444, NT2RP3000481, NT2RP3000616, NT2RP3000871, NT2RP3001012, NT2RP3001061, NT2RP3001240, NT2RP3001322, NT2RP3001542, NT2RP3002286, NT2RP3002342, NT2RP3002448, NT2RP3002571, NT2RP3002664, NT2RP3002790, NT2RP3002887, NT2RP3002983, NT2RP3003527, NT2RP3003535, NT2RP3003559, NT2RP3004000, NT2RP3004075, NT2RP3004345, NT2RP4000962, NT2RP4001001, NT2RP4001009, NT2RP4001467, NT2RP4002451, OVARC1000003, OVARC1000275, OVARC1000298, OVARC1000307, OVARC1000811, OVARC1001132, OVARC1001222, OVARC1001338, OVARC1001607, OVARC1001725, OVARC1001727, OVARC1002058, OVARC1002178, PLACE1000033, PLACE1000740, PLACE1000914, PLACE1000986, PLACE1001123, PLACE1001231, PLACE1001401, PLACE1001464, PLACE1001536, PLACE1001564, PLACE1001788, PLACE1001795, PLACE1001949, PLACE1002355, PLACE1003135, PLACE1003573, PLACE1003737, PLACE1004028, PLACE1004450, PLACE1004519, PLACE1005003, PLACE1005239, PLACE1005250, PLACE1005519, PLACE1005601, PLACE1005660, PLACE1005669, PLACE1005682, PLACE1005745, PLACE1006093, PLACE1006208, PLACE1006219, PLACE1006290, PLACE1006515, PLACE1006786, PLACE1006959, PLACE1007028, PLACE1007040, PLACE1007081, PLACE1007296, PLACE1007591, PLACE1007845, PLACE1007881, PLACE1007971, PLACE1008282, PLACE1008297, PLACE1008359, PLACE1008549, PLACE1008657, PLACE1008716, PLACE1008744, PLACE1008984, PLACE1009196, PLACE1009279, PLACE1009546, PLACE1009600, PLACE1009735, PLACE1010011, PLACE1010081, PLACE1010713, PLACE1010784, PLACE1010827, PLACE1010968, PLACE1011407, PLACE1011824, PLACE3000213, SKNMC1000004, SKNMC1000082, THYRO1000036, THYRO1000196, THYRO1000400, THYRO1000580, THYRO1000678, THYRO1000776, THYRO1000846, THYRO1000956, THYRO1001071, THYRO1001102, THYRO1001266, THYRO1001327, THYRO1001471, Y79AA1000876, Y79AA1000959, Y79AA1001056, Y79AA1001062, Y79AA1001264, Y79AA1001795.

**[0209]** Genes exhibiting characteristic features in the expression thereof were selected by statistical analysis of these

data. Two examples are shown below to describe the selection of genes of which expression is varied greatly among tissues. The  $\beta$ -actin gene is used frequently as a control in gene expression analysis. Genes of which expression is varied greatly among tissues as compared that of the  $\beta$ -actin gene were determined as follows. Specifically, sum of squared deviation was calculated in the signal intensity of  $\beta$ -actin observed in each tissue, which was divided by 7

degrees of freedom to determine a variance  $S_a^2$ . Next, sum of squared deviation was calculated in the signal intensity of a compared gene in each tissue, which was divided by 7 degrees of freedom to determine a variance  $S_b^2$ . By taking variance ratio  $F$  as  $F=S_b^2/S_a^2$ , genes with a significance level of 5% or more were extracted in the  $F$  distribution. Genes extracted are indicated below by the corresponding clone names: clone: BNGH41000020, NT2RM4000761, Y79AA1002376.

**[0210]** Gene of OVARC1000037(heterogeneous nuclear ribonucleoprotein (hnRNP)) which expression is varied little. Genes of which expression is varied greatly among tissues as compared that of the OVARC1000037 gene were determined as follows. Specifically, sum of squared deviation was calculated in the signal intensity of  $\beta$ -actin observed in each tissue, which was divided by 7 degrees of freedom to determine a variance  $S_a^2$ . Next, sum of squared deviation was calculated in the signal intensity of a gene to be compared observed in each tissue, which was divided by 7 degrees of freedom to determine a variance  $S_b^2$ . By taking variance ratio  $F$  as  $F=S_b^2/S_a^2$ , genes with a significance level of 5% or more were extracted in the  $F$  distribution. Genes extracted are indicated below by the corresponding clone names: clone: BNGH41000020, HEMBA1000300, OVARC1001030, NT2RM4000761, PLACE1000231, HEMBA1002316, NT2RP1000325, NT2RP1000271, PLACE1004648, HEMBA1005145, HEMBA1005929, NT2RP2002510, NT2RP2001538, NT2RP2002409, NT2RP2002188, NT2RP2001903, NT2RP2002533, NT2RP2002015, NT2RP2006269, NT2RP2004837, NT2RP2004205, NT2RP2005378, HEMBA1006357, HEMBB1000567, NT2RP2003940, NT2RP2004794, HEMBA1006912, NT2RP2004556, NT2RP2005163, NT2RP3000838, NT2RP3001271, PLACE2000118, NT2RP3000645, NT2RP3003076, HEMBB1002693, MAMMA1000046, NT2RP3003354, THYR01001205, MAMMA1000614, MAMMA1000652, MAMMA1000810, THYRO1001242, MAMMA1001066, MAMMA1002224, MAMMA1001634, MAMMA1001094, MAMMA1002205, NT2RM1000855, NT2RM1000858, Y79AA1002376, NT2RM2000423.

**[0211]** Thus, characteristic features in the expression of a gene are illustrated by comparing and statistically analyzing the expression of many genes.

#### Analysis of disease-associated genes

**[0212]** Non-enzymic protein glycation reaction is believed to be a cause of a variety of chronic diabetic complications. Accordingly, genes of which expression is elevated or decreased in a glycosylated protein-specific manner in the endothelial cells are associated with diabetic complications caused by glycosylated proteins. Vascular endothelial cells are affected with glycosylated proteins present in blood. Reaction products of non-enzymic protein glycation include amadori compound (glycosylated protein) as a mildly glycosylated protein and advanced glycation endproduct as a heavily glycosylated protein. Hence, a survey was carried out for genes of which expression levels are varied depending on the presence of these glycosylated proteins in endothelial cells. The mRNAs were extracted from endothelial cells that were cultured in the presence or absence of glycosylated protein. The mRNAs were converted into radiolabeled first strand cDNAs for preparing probes. The probes were hybridized to the above-mentioned DNA array. Signal of each DNA spot was detected by BAS2000 and analyzed by ArrayGauge (Fuji Photo Film Co., Ltd.).

**[0213]** Advanced glycation endproduct of bovine serum albumin was prepared as follows: bovine serum albumin (BSA; Sigma) was incubated in a phosphate buffer solution containing 50 mM glucose at 37 for 8 weeks; and the resulting brownish BSA was dialyzed against a phosphate buffer solution.

**[0214]** Human normal pulmonary arterial endothelial cells (Cell Applications) were cultured in an Endothelial Cell Growth Medium (Cell Applications). The culture dish (Falcon) with the cells were incubated in a CO<sub>2</sub> incubator (37., 5% CO<sub>2</sub>, in a humid atmosphere). When the cells were grown to be confluent in the dish, 250.g/ml of bovine serum albumin (sigma), glycosylated bovine serum albumin (Sigma) or advanced glycation endproduct of bovine serum albumin was added thereto and the cells were incubated for 33 hours. The mRNA was extracted from the cells by using a FastTrack<sup>TM</sup> 2.0 kit (Invitrogen). The labeling of hybridization probe was carried out by using the mRNA according to the same procedure as described above.

**[0215]** Table 185 shows the expression level of each cDNA in human pulmonary arterial endothelial cells cultured in a medium containing bovine serum albumin (sigma), glycosylated bovine serum albumin (Sigma) or advanced glycation endproduct of bovine serum albumin. Genes of which expression was detected in the endothelial cell are as follows: BNGH41000020, BNGH41000087, HEMBA1000275, HEMBA1000300, HEMBA1000477, HEMBA1000634, HEMBA1000671, HEMBA1000713, HEMBA1000745, HEMBA1000835, HEMBA1000875, HEMBA1000940, HEMBA1001390, HEMBA1002131, HEMBA1002163, HEMBA1002164, HEMBA1002195, HEMBA1002227, HEMBA1002239, HEMBA1002420, HEMBA1002767, HEMBA1002992, HEMBA1003047, HEMBA1003120, HEMBA1003294, HEMBA1003315, HEMBA1003602, HEMBA1003945, HEMBA1004007, HEMBA1004067,



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	HEMBA1004971,	HEMBA1005145,	HEMBA1005267,	HEMBA1005337,	HEMBA1005698,	HEMBA1005929,
	HEMBA1005945,	HEMBA1006171,	HEMBA1006299,	HEMBA1006335,	HEMBA1006357,	HEMBA1006430,
	HEMBA1006482,	HEMBA1006658,	HEMBA1006724,	HEMBA1006770,	HEMBA1006912,	HEMBA1006960,
	HEMBA1007063,	HEMBB1000447,	HEMBB1000642,	HEMBB1000905,	HEMBB1001026,	HEMBB1001048,
5	HEMBB1001573,	HEMBB1001847,	HEMBB1001978,	HEMBB1002041,	HEMBB1002427,	HEMBB1002663,
	HEMBB1002693,	MAMMA1000102,	MAMMA1000106,	MAMMA1000204,	MAMMA1000403,	MAMMA1000449,
	MAMMA1000614,	MAMMA1000652,	MAMMA1000810,	MAMMA1000814,	MAMMA1000881,	MAMMA1000986,
	MAMMA1001066,	MAMMA1001237,	MAMMA1001284,	MAMMA1001344,	MAMMA1001615,	MAMMA1001634,
	MAMMA1001893,	MAMMA1001901,	MAMMA1001957,	MAMMA1002087,	MAMMA1002095,	MAMMA1002165,
10	MAMMA1002205,	MAMMA1002224,	MAMMA1002633,	MAMMA1003126,	NT2RM1000462,	NT2RM1000580,
	NT2RM1000789,	NT2RM1000855,	NT2RM1000858,			
	NT2RM2000241,	NT2RM2000306,	NT2RM2000410,	NT2RM2000423,	NT2RM2000582,	NT2RM2000589,
	NT2RM2000622,	NT2RM2000773,	NT2RM4000100,	NT2RM4000198,	NT2RM4000284,	NT2RM4000444,
	NT2RM4000587,	NT2RM4000593,	NT2RM4000761,	NT2RM4000997,	NT2RM4001321,	NT2RM4001325,
15	NT2RM4001377,	NT2RM4001735,	NT2RM4001768,	NT2RM4001843,	NT2RP1000002,	NT2RP1000181,
	NT2RP1000271,	NT2RP1000300,	NT2RP1000325,	NT2RP1000465,	NT2RP1000740,	NT2RP1000981,
	NT2RP2000092,	NT2RP2000240,	NT2RP2000479,	NT2RP2000533,	NT2RP2000610,	NT2RP2000616,
	NT2RP2000649,	NT2RP2000663,	NT2RP2000712,	NT2RP2000903,	NT2RP2001276,	NT2RP2001388,
	NT2RP2001480,	NT2RP2001495,	NT2RP2001529,	NT2RP2001538,	NT2RP2001662,	NT2RP2001878,
20	NT2RP2001903,	NT2RP2001948,	NT2RP2001956,	NT2RP2002015,	NT2RP2002188,	NT2RP2002232,
	NT2RP2002409,	NT2RP2002510,	NT2RP2002527,	NT2RP2002533,	NT2RP2002564,	NT2RP2002721,
	NT2RP2002824,	NT2RP2002942,	NT2RP2002976,	NT2RP2003138,	NT2RP2003210,	NT2RP2003390,
	NT2RP2003593,	NT2RP2003599,	NT2RP2003664,	NT2RP2003931,	NT2RP2003940,	NT2RP2004069,
	NT2RP2004108,	NT2RP2004179,	NT2RP2004205,	NT2RP2004495,	NT2RP2004524,	NT2RP2004556,
25	NT2RP2004606,	NT2RP2004648,	NT2RP2004794,	NT2RP2004837,	NT2RP2004847,	NT2RP2005027,
	NT2RP2005069,	NT2RP2005163,	NT2RP2005247,	NT2RP2005378,	NT2RP2005425,	NT2RP2005535,
	NT2RP2005541,	NT2RP2005632,	NT2RP2005774,			
	NT2RP2005878,	NT2RP2006099,	NT2RP2006134,	NT2RP2006269,	NT2RP2006512,	NT2RP3000011,
	NT2RP3000171,	NT2RP3000201,	NT2RP3000232,	NT2RP3000436,	NT2RP3000460,	NT2RP3000645,
30	NT2RP3000652,	NT2RP3000676,	NT2RP3000721,	NT2RP3000818,	NT2RP3000820,	NT2RP3000838,
	NT2RP3000907,	NT2RP3001159,	NT2RP3001195,	NT2RP3001240,	NT2RP3001271,	NT2RP3001388,
	NT2RP3001592,	NT2RP3001738,	NT2RP3001754,	NT2RP3002015,	NT2RP3002324,	NT2RP3002342,
	NT2RP3002353,	NT2RP3002409,	NT2RP3002448,	NT2RP3002721,	NT2RP3002737,	NT2RP3002738,
	NT2RP3002836,	NT2RP3002900,	NT2RP3003076,	NT2RP3003354,	NT2RP3003448,	NT2RP3003473,
35	NT2RP3003532,	NT2RP3003614,	NT2RP3003939,	NT2RP3003963,	NT2RP3004025,	NT2RP3004067,
	NT2RP3004075,	NT2RP3004083,	NT2RP3004090,	NT2RP3004119,	NT2RP3004130,	NT2RP3004133,
	NT2RP3004294,	NT2RP3004309,	NT2RP3004345,	NT2RP3004374,	NT2RP3004557,	NT2RP3004625,
	NT2RP3004640,	NT2RP3004647,	NT2RP4000108,	NT2RP4000634,	NT2RP4001001,	NT2RP4001009,
	NT2RP4001467,	NT2RP4001877,	NT2RP4001879,	NT2RP4002187,	NT2RP4002451,	NT2RP4002715,
40	OVARC1000003,	OVARC1000090,	OVARC1000105,	OVARC1000137,	OVARC1000208,	OVARC1000298,
	OVARC1000313,	OVARC1000331,	OVARC1000410,	OVARC1000439,	OVARC1000553,	OVARC1000775,
	OVARC1000853,	OVARC1000873,	OVARC1000916,	OVARC1000956,	OVARC1000995,	OVARC1001030,
	OVARC1001049,	OVARC1001086,	OVARC1001132,			
	OVARC1001222,	OVARC1001260,	OVARC1001336,	OVARC1001569,	OVARC1001570,	OVARC1001596,
45	OVARC1001607,	OVARC1001807,	OVARC1001991,	PLACE1000231,	PLACE1000258,	PLACE1000442,
	PLACE1000740,	PLACE1000927,	PLACE1001016,	PLACE1001100,	PLACE1001114,	PLACE1001123,
	PLACE1001229,	PLACE1001340,	PLACE1001407,	PLACE1001464,	PLACE1001788,	PLACE1001795,
	PLACE1001918,	PLACE1002080,	PLACE1002095,	PLACE1002329,	PLACE1002374,	PLACE1002518,
	PLACE1002547,	PLACE1002726,	PLACE1002905,	PLACE1002911,	PLACE1002967,	PLACE1003163,
50	PLACE1003407,	PLACE1003460,	PLACE1003573,	PLACE1003598,	PLACE1003644,	PLACE1003772,
	PLACE1003839,	PLACE1003845,	PLACE1004078,	PLACE1004166,	PLACE1004168,	PLACE1004199,
	PLACE1004279,	PLACE1004282,	PLACE1004441,	PLACE1004482,	PLACE1004492,	PLACE1004637,
	PLACE1004887,	PLACE1005003,	PLACE1005005,	PLACE1005031,	PLACE1005250,	PLACE1005410,
	PLACE1005519,	PLACE1005544,	PLACE1005660,	PLACE1005669,	PLACE1005725,	PLACE1005736,
55	PLACE1005745,	PLACE1005768,	PLACE1005815,	PLACE1006073,	PLACE1006208,	PLACE1006219,
	PLACE1006290,	PLACE1006443,	PLACE1006809,	PLACE1006959,	PLACE1007028,	PLACE1007296,
	PLACE1007626,	PLACE1007702,	PLACE1007845,	PLACE1008282,	PLACE1008469,	PLACE1008657,
	PLACE1009196,	PLACE1009600,	PLACE1003735,	PLACE1010081,	PLACE1010251,	PLACE1010713,

PLACE1011116, PLACE1011181,  
 PLACE1011236, PLACE1011516, PLACE1011708, PLACE1011824, PLACE1011978, PLACE2000118,  
 PLACE3000181, SKNMC1000004, SKNMC1000014, THYRO1000584, THYRO1000866, THYRO1001113,  
 THYRO1001128, THYRO1001205, THYRO1001242, THYRO1001495, THYRO1001523, THYRO1001529,  
 5 THYRO1001593, THYRO1001608, THYRO1001702, THYRO1001725, THYRO1001770, THYRO1001803,  
 Y79AA1000117, Y79AA1000207, Y79AA1000226, Y79AA1000270, Y79AA1000426, Y79AA1000777,  
 Y79AA1000876, Y79AA1000888, Y79AA1000959, Y79AA1001013, Y79AA1001056, Y79AA1001090,  
 Y79AA1001264, Y79AA1001272, Y79AA1001328, Y79AA1001427, Y79AA1001430, Y79AA1001530,  
 Y79AA1001592, Y79AA1001727, Y79AA1001793, Y79AA1001799, Y79AA1001863, Y79AA1002022,  
 10 Y79AA1002213, Y79AA1002373, Y79AA1002376, Y79AA1002381.

[0216] Signal ratios of EC\_AGE\_BSA to EC\_BSA and of EC\_glycated\_BSA to EC\_BSA were calculated for each gene. Genes with high signal ratios were selected. In the case of calculating the ratio of signal value of 40 or less to that of more than 40, such signal values were, for convenience, taken as 40 instead of the real values. When the ratio EC\_AGE\_BSA/EC\_BSA is 2 or more, expression of the genes exhibiting such ratio is expected to be elevated due to advanced glycation endproduct of bovine serum albumin. The higher the value is, the higher the gene expression level is. When the ratio EC\_AGE\_BSA/EC\_BSA ranges from 0.5 to 2, expression of the genes exhibiting such ratio is expected to be unaffected due to advanced glycation endproduct of bovine serum albumin. When the ratio EC\_AGE\_BSA/EC\_BSA is less than 0.5, expression of the genes exhibiting such ratio value is expected to be decreased due to advanced glycation endproduct of bovine serum albumin. The lower the value is, the lower the gene expression level is.

20 [0217] Clone with EC\_AGE\_BSA/EC\_BSA ratio of 2 or higher are as follows: NT2RP2001538, NT2RP4001001 and Y79AA1000967.

[0218] These cDNAs are associated with diabetes.

#### Analysis of genes associated with neural cell differentiation

25 [0219] Genes involved in neural cell differentiation are useful for treating neurological diseases. It is possible that genes with varying expression levels in response to induction of cellular differentiation in neural cells are associated with neurological diseases.

30 [0220] A survey was performed for genes of which expression levels are varied in response to induction of differentiation (stimulation by retinoic acid (RA)) in cultured cells of a neural strain, NT2.

[0221] The NT2 cells were treated basically according to supplier's instruction manual. "Undifferentiated NT2 cells" means NT2 cells successively cultured in an Opti-MEM I (GIBCO-BRL; catalog No. 31985) containing 10%(v/v) fetal bovine serum and 1%(v/v) penicillin-streptomycin (GIBCO BRL). "NT2 cells cultured in the presence of retinoic acid" means the cells resulted from transferring undifferentiated NT2 cells into a retinoic acid-containing medium, which consists of D-MEM (GIBCO BRL; catalog No. 11965), 10%(v/v) fetal bovine serum, 1%(v/v) penicillin-streptomycin and 10.M retinoic acid (GIBCO-BRL), and the subsequent successive culture therein for 5 weeks. "NT2 cells that were cultured in the presence of retinoic acid and then further cultured in the presence of cell-division inhibitor added" means NT2 cells resulted from transferring NT2 cells cultured in the presence of retinoic acid for 5 weeks into a cell-division inhibitor-containing medium, which consisted of D-MEM(GIBCO BRL; catalog No.11965), 10%(v/v) fetal bovine serum, 1%(v/v) penicillin-streptomycin, 10. M retinoic acid, 10.M FudR (5-fluoro-2'-deoxyuridine: GIBCO BRL), 10. M Urd (Uridine: GIBCO BRL) and 1.M araC (Cytosine.-D-Arabinofuranoside: GIBCO BRL), and the subsequent successive culture for 2 weeks. Each of the cells were treated with trypsin and then harvested. Total RNAs were extracted from the cells by using S.N.A.P.<sup>(TM)</sup> Total RNA Isolation kit (Invitrogen). The labeling of probe used for hybridization was carried out by using 10.g of the total RNA according to the same methods as described above. The data were obtained in triplicate (n=3). The data of signal value representing gene expression level in the cells in the presence of stimulation for inducing differentiation were compared with those in the absence of the stimulation. The comparison was performed by statistical treatment of two-sample t-test. Clones with significant difference in the signal distribution were selected under the condition of p<0.05. In this analysis, clones with the difference can be statistically detected even when the signals were low. Accordingly, clones with signal value of 40 or less were also assessed for the selection.

50 [0222] Tables 186-365 show the expression level of each cDNA in undifferentiated NT2 cells, NT2 cells cultured in the presence of RA, and NT2 cells that were cultured in the presence of RA and that were further cultured in the presence of cell-division inhibitor added.

[0223] Averaged signal values ( $M_1$ ,  $M_2$ ) and sample variances ( $s_1^2$ ,  $s_2^2$ ) were calculated for each gene in each of the cells, and then, the pooled sample variances  $s^2$  were obtained from the sample variances of the two types of cells to be compared. The t values were determined according to the following formula:  $t=(M_1-M_2)/s/(1/3+1/3)^{1/2}$ . When the determined t-value was greater than a t-value at P, which means the probability of significance level, of 0.05 or 0.01 in the t-distribution table with 4 degrees of freedom, the difference was judged to be found in the expression level of the gene between the two types of cells at p<0.05 or p<0.01, respectively. The tables also include the information on

an increase (+) or decrease (-) in the expression level of a gene in the treated cells when the level is compared with that of untreated undifferentiated cells.

**[0224]** Clones of which expression levels increased by RA are as follows: HEMBA1000121, HEMBA1000275, HEMBA1000300, HEMBA1000634, HEMBA1000671, HEMBA1000875, HEMBA1001184, HEMBA1001390, HEMBA1001886, HEMBA1002163, HEMBA1002227, HEMBA1002420, HEMBA1002421, HEMBA1003072, HEMBA1003120, HEMBA1003294, HEMBA1003497, HEMBA1004007, HEMBA1004110, HEMBA1004391, HEMBA1004444, HEMBA1005230, HEMBA1005246, HEMBA1005267, HEMBA1005489, HEMBA1005913, HEMBA1006299, HEMBA1006357, HEMBA1006517, HEMBA1006544, HEMBA1006658, HEMBA1006749, HEMBA1007063, HEMBA1007241, HEMBB1000447, HEMBB1000542, HEMBB1000567, HEMBB1000642, HEMBB1000668, HEMBB1001026, HEMBB1001847, HEMBB1002051, HEMBB1002120, HEMBB1002228, HEMBB1002693, MAMMA1000106, MAMMA1000141, MAMMA1000473, MAMMA1000528, MAMMA1000810, MAMMA1000881, MAMMA1001634, MAMMA1001957, MAMMA1002205, MAMMA1002224, NT2RM2000423, NT2RM2000497, NT2RM2000582, NT2RM2001126, NT2RM2001902, NT2RM4000198, NT2RM4000284, NT2RM4000593, NT2RM4001321, NT2RP1000002, NT2RP1000050, NT2RP1000181, NT2RP1000261, NT2RP1000465, NT2RP1000468, NT2RP1000579, NT2RP1000679, NT2RP2000092, NT2RP2000479, NT2RP2000610, NT2RP2000663, NT2RP2000694, NT2RP2000903, NT2RP2001388, NT2RP2001538, NT2RP2001878, NT2RP2001015, NT2RP2002304, NT2RP2002721, NT2RP2002824, NT2RP2002942, NT2RP2002974, NT2RP2002976, NT2RP2003179, NT2RP2003302, NT2RP2003383, NT2RP2003469, NT2RP2003664, NT2RP2003940, NT2RP2004069, NT2RP2004108, NT2RP2004524, NT2RP2004556, NT2RP2004670, NT2RP2005069, NT2RP2005247, NT2RP2005425, NT2RP2005463, NT2RP2005514, NT2RP2005535, NT2RP2005541, NT2RP2005774, NT2RP2005878, NT2RP2005883, NT2RP2005887, NT2RP2006099, NT2RP2006134, NT2RP3000011, NT2RP3000125, NT2RP3000171, NT2RP3000232, NT2RP3000460, NT2RP3000481, NT2RP3000652, NT2RP3000677, NT2RP3000818, NT2RP3000820, NT2RP3001044, NT2RP3001061, NT2RP3001170, NT2RP3001240, NT2RP3001322, NT2RP3001388, NT2RP3001542, NT2RP3001592, NT2RP3001976, NT2RP3002790, NT2RP3002900, NT2RP3002983, NT2RP3003000, NT2RP3003354, NT2RP3003532, NT2RP3003729, NT2RP3003874, NT2RP3003939, NT2RP3004025, NT2RP3004083, NT2RP3004090, NT2RP3004130, NT2RP3004202, NT2RP3004294, NT2RP3004640, NT2RP4000108, NT2RP4000634, NT2RP4002451, NT2RP4002715, OVARC1000090, OVARC1000208, OVARC1000275, OVARC1000553, OVARC1000775, OVARC1000853, OVARC1000873, OVARC1000916, OVARC1000995, OVARC1001030, OVARC1001049, OVARC1001132, OVARC1001596, OVARC1002178, PLACE1000258, PLACE1000442, PLACE1000927, PLACE1000986, PLACE1001100, PLACE1001123, PLACE1001795, PLACE1002518, PLACE1002547, PLACE1002967, PLACE1003407, PLACE1003428, PLACE1003644, PLACE1003839, PLACE1004078, PLACE1004441, PLACE1004450, PLACE1005669, PLACE1005682, PLACE1005736, PLACE1005768, PLACE1005815, PLACE1006073, PLACE1006208, PLACE1007296, PLACE1007626, PLACE1008282, PLACE1008984, PLACE1008985, PLACE1010445, PLACE1011708, PLACE1011978, PLACE4000455, SKNMC1000004, THYRO1000036, THYRO1000580, THYRO1000776, THYRO1000999, THYRO1001063, THYRO1001128, THYRO1001205, THYRO1001327, THYRO1001523, THYRO1001725, THYRO1001770, Y79AA1000207, Y79AA1000226, Y79AA1000270, Y79AA1001056, Y79AA1001062, Y79AA1001090, Y79AA1001727, Y79AA1002213, Y79AA1002381.

**[0225]** Clones of which expression levels decreased by RA are as follows: BNGH41000020, HEMBA1005070, NT2RP2005027, NT2RP3003473, Y79AA1002376.

**[0226]** Clones of which expression levels increase by RA/inhibitor are as follows:

HEMBA1000128, HEMBA1000875, HEMBA1001390, HEMBA1002163, HEMBA1002227, HEMBA1002421, HEMBA1004391, HEMBA1004454, HEMBA1004785, HEMBA1005913, HEMBA1006171, HEMBA1006299, HEMBA1006335, HEMBA1006544, HEMBA1007241, HEMBB1000447, HEMBB1000668, MAMMA1000994, MAMMA1001344, NT2RM2000582, NT2RP1001004, NT2RP2000663, NT2RP2000694, NT2RP2000903, NT2RP2001388, NT2RP2002674, NT2RP2002974, NT2RP2003383, NT2RP2004069, NT2RP2004606, NT2RP2004837, NT2RP2005069, NT2RP2005425, NT2RP2005463, NT2RP2005541, NT2RP2005883, NT2RP2005887, NT2RP3000460, NT2RP3000838, NT2RP3001044, NT2RP3001240, NT2RP3001388, NT2RP3002721, NT2RP3002738, NT2RP3003469, NT2RP3004083, NT2RP3004130, NT2RP3004202, NT2RP3004294, NT2RP3004640, NT2RP4000108, NT2RP4002451, NT2RP4002715, OVARC1000275, OVARC1000467, OVARC1000553, OVARC1000853, OVARC1000873, OVARC1000916, OVARC1000995, OVARC1001030, OVARC1001222, OVARC1001596, OVARC1002058, OVARC1002178, PLACE1000927, PLACE1001123, PLACE1001407, PLACE1001464, PLACE1001564, PLACE1001795, PLACE1002547, PLACE1003407, PLACE1003644, PLACE1003845, PLACE1004441, PLACE1004482, PLACE1005410, PLACE1005601, PLACE1005725, PLACE1005736, PLACE1006093, PLACE1006219, PLACE1006290, PLACE1006716, PLACE1007296, PLACE1007626, PLACE1008359, PLACE1010968, PLACE1011364,

PLACE1011824, THYRO1000678, THYRO1000776, THYRO1000999, THYRO1001113, THYRO1001237, THYRO1001523, Y79AA1000226, Y79AA1000888, Y79AA1001430.

[0227] Clones of which expression levels decrease by RA/inhibitor are as follows: HEMBA1000349, HEMBA1001297, HEMBA1001878, HEMBA1005070, HEMBA1006482, HEMBB1001959, NT2RM2001939, NT2RP1000981, NT2RP2001469, NT2RP3003473, OVARC1001132, PLACE1001655, Y79AA1000127, Y79AA1002381.

[0228] Clones of which expression levels increase in the presence of both RA and RA/inhibitor are as follows: HEMBA1000875, HEMBA1001390, HEMBA1002163, HEMBA1002227, HEMBA1002421, HEMBA1004391, HEMBA1005913, HEMBA1006299, HEMBA1006544, HEMBA1007241, HEMBB1000447, HEMBB1000668, NT2RM2000582, NT2RP2000663, NT2RP2000694, NT2RP2000903, NT2RP2001388, NT2RP2002974, NT2RP2003383, NT2RP2004069, NT2RP2005069, NT2RP2005425, NT2RP2005463, NT2RP2005541, NT2RP2005883, NT2RP2005887, NT2RP3000460, NT2RP3001044, NT2RP3001240, NT2RP3001388, NT2RP3004083, NT2RP3004130, NT2RP3004202, NT2RP3004294, NT2RP3004640, NT2RP4000108, NT2RP4002451, NT2RP4002715, OVARC1000275, OVARC1000553, OVARC1000853, OVARC1000873, OVARC1000916, OVARC1000995, OVARC1001030, OVARC1001596, OVARC1002178, PLACE1000927, PLACE1001123, PLACE1001795, PLACE1002547, PLACE1003407, PLACE1003644, PLACE1004441, PLACE1005736, PLACE1007296, PLACE1007626, THYRO1000776, THYRO1000999, THYRO1001523, Y79AA1000226.

[0229] Clones of which expression levels decrease in the presence of both RA and RA/inhibitor are as follows: HEMBA1005070 and NT2RP3003473.

[0230] These are neurological disease-associated clones.

#### Analysis of rheumatoid arthritis-associated genes

[0231] The onset of rheumatoid arthritis is thought to be involved in the proliferation of synovial cells covering inner surfaces of joint cavity and in inflammatory reaction resulted from the action of cytokines produced by leukocytes infiltrating into the joint synovial tissues (Rheumatism Information Center.<http://www.rheuma-net.or.jp/>). Recent studies have also revealed that tissue necrosis factor (TNF)- $\alpha$  participates in the onset (Current opinion in immunology 1999, 11, 657-662). When the expression of a gene exhibits responsiveness to the action of TNF on synovial cells, the gene is considered to be involved in rheumatoid arthritis.

[0232] A survey was performed for genes of which expression levels are varied in response to TNF- $\alpha$  in the primary cell culture of synovial tissue. The primary cultured cells of the smooth muscle (Cell Applications) were grown to be confluent in a culture dish, and then, human TNF- $\alpha$  (Boehringer-Mannheim) was added at a final concentration of 10 ng/ml thereto. The culture was further continued for 24 hours.

[0233] Total RNA was extracted from the cells by using S.N.A.P.<sup>(TM)</sup> Total RNA Isolation kit (Invitrogen). The labeling of probe used for hybridization was carried out by using 10 $\mu$ g of the total RNA according to the same methods as described above. The data were obtained in triplicate (n=3). The data of signal value representing gene expression level in the cells in the presence of TNF stimulation were compared with those in the absence of the stimulation. The comparison was performed by statistical treatment of two-sample t-test. Clones with significant difference in the signal distribution were selected under the condition of p<0.05. In this analysis, clones with the difference can be statistically detected even when the signals were low. Accordingly, clones with signal value of 40 or less were also assessed for the selection.

[0234] Table 366 shows the expression level of each cDNA in synovial cells cultured in the absence or presence of TNF.

[0235] Averaged signal values ( $M_1$ ,  $M_2$ ) and sample variances ( $s_1^2$ ,  $s_2^2$ ) for each gene were calculated in each of the cells, and then, the pooled sample variances  $s^2$  were obtained from the sample variances of the two types of cells to be compared. The t-values were determined according to the following formula:  $t=(M_1-M_2)/s/(1/3+1/3)^{1/2}$ . When the determined t-value was greater than a t-value at P, which means the probability of significance level, of 0.05 or 0.01 in the t-distribution table with 4 degrees of freedom, the difference was judged to be found in the expression level of the gene between the two types of cells at p<0.05 or p<0.01, respectively. The tables also include the information of an increase (+) or decrease (-) in the expression level of a gene in the stimulated cells when the level is compared with that of unstimulated cells.

[0236] Clones of which expression levels are elevated by TNF- $\alpha$  are as follows:

BNGH41000020, HEMBA1000349, HEMBA1000634, HEMBA1000671, HEMBA1000835, HEMBA1000962, HEMBA1002178, HEMBA1002195, HEMBA1002239, HEMBA1002420, HEMBA1002524, HEMBA1002992, HEMBA1003315, HEMBA1003392, HEMBA1003487, HEMBA1003602, HEMBA1004067, HEMBA1004797, HEMBA1005337, HEMBA1005489, HEMBA1006916, HEMBB1000668, HEMBB1000905, HEMBB1001547, HEMBB1001573, HEMBB1002041, HEMBB1002663, MAMMA1000652, MAMMA1000810, MAMMA1001634,

MAMMA1002091, MAMMA1002234, NT2RM2000306, NT2RM4000417, NT2RP1000002, NT2RP1000181,  
 NT2RP1000740, NT2RP2000694, NT2RP2001921, NT2RP2002527, NT2RP2004495, NT2RP2004606,  
 NT2RP2005163, NT2RP2005463, NT2RP2006134, NT2RP3000171, NT2RP3000652, NT2RP3001195,  
 NT2RP3001976, NT2RP3003473, NT2RP3003874, NT2RP3004090, NT2RP3004294, NT2RP3004557,  
 5 NT2RP3004647, NT2RP4000108, NT2RP4001001, NT2RP4001877, OVARC1000090, OVARC1000105,  
 OVARC1000275, OVARC1000439, OVARC1001607, PLACE1000740, PLACE1000927, PLACE1001016,  
 PLACE1001100, PLACE1001464, PLACE1001500, PLACE1001918, PLACE1002095, PLACE1002547,  
 PLACE1003644, PLACE1004519, PLACE1005031, PLACE1005410, PLACE1005736, PLACE1006219,  
 PLACE1006809, PLACE1008716, PLACE1010081, THYRO1001770, Y79AA1000127, Y79AA1000207,  
 10 Y79AA1000270, Y79AA1000876, Y79AA1001013, Y79AA1001264, Y79AA1001272, Y79AA1001328,  
 Y79AA1001430, Y79AA1001530, Y79AA1001799.

[0237] Clones of which expression levels decrease by TNF- are as follows:

NT2RM4000326, NT2RP1000300, NT2RP2000514, NT2RP2001755, NT2RP2006042, NT2RP3000481,  
 NT2RP3002790. These are rheumatoid arthritis-associated clones.

#### EXAMPLE 16

Search for a signal sequence, transmembrane region and functional domain in deduced amino acid sequences

20 [0238] The deduced amino acid sequences from the full-length nucleotide sequences were examined to predict the  
 presence of a signal sequence in their amino-termini as well as the presence of a transmembrane region. The amino  
 acid sequences were also searched for a protein functional domain (motif). The examinations for a signal sequence  
 in the amino-terminus, for a transmembrane region and for a functional domain were performed by using PSORT [K.  
 Nakai & M. Kanehisa, Genomics, 14:897-911 (1992)], SOSUI [T. Hirokawa et al., Bioinformatics, 14:378-379 (1998)]  
 25 (Mitsui Knowledge Industry Co., Ltd.) and Pfam (<http://www.sanger.ac.uk/Software/Pfam/index.shtml>), respectively.  
 When the presence of a signal sequence or a transmembrane region in the amino-terminus was predicted in the amino  
 acid sequence by PSORT or SOSUI, the protein was predicted to be a secretory protein or a transmembrane protein.  
 When the amino acid sequence matched a functional domain in the Pfam search for a functional domain, the function  
 of the protein is predictable based on the matching data, for example, by referring to the functional categories in  
 30 PROSITE (<http://www.expasy.ch/cgi-bin/prosite-list.pl>). The functional domain search can be performed by using  
 PROSITE instead of Pfam.

[0239] Search results obtained by using the respective software programs are indicated below.

[0240] Clones whose deduced amino acid sequences were predicted to have signal sequences by PSORT search  
 are as follows:

35 HEMBA1000713, HEMBA1002420, HEMBA1002421, HEMBA1003101, HEMBA1004110, HEMBA1006707,  
 HEMBA1006902, HEMBB1001530, HEMBB1001573, HEMBB1001978, HEMBB1002162, HEMBB1002245,  
 HEMBB1002427, MAMMA1000102, MAMMA1000118, MAMMA1000457, MAMMA1001043, MAMMA1001344,  
 MAMMA1001893, MAMMA1002070, MAMMA1002165, MAMMA1002633, NT2RM2000241, NT2RM2000410,  
 NT2RM2001941, NT2RM4001325, NT2RP1001563, NT2RP2001495, NT2RP2002063, NT2RP2002721,  
 40 NT2RP2003383, NT2RP2003593, NT2RP2003655, NT2RP2003664, NT2RP2004179, NT2RP2004205,  
 NT2RP2004524, NT2RP2005463, NT2RP3000460, NT2RP3001012, NT2RP3001858, NT2RP3002836,  
 NT2RP3003076, NT2RP3003532, NT2RP3004133, NT2RP3004309, NT2RP4001467, NT2RP4002451,  
 OVARC1000298, OVARC1000811, PLACE1000231, PLACE1000740, PLACE1001183, PLACE1001536,  
 PLACE1001564, PLACE1002095, PLACE1002374, PLACE1003839, PLACE1001482, PLACE1005005,  
 45 PLACE1005250, PLACE1005383, PLACE1005410, PLACE1005544, PLACE1005569, PLACE1006093,  
 PLACE1006277, PLACE1006809, PLACE1007626, PLACE1008359, PLACE1009067, PLACE1010251,  
 PLACE1011236, SKNMC1000004, SKNMC1000014, THYRO1000099, THYRO1000196, THYRO1001237,  
 THYRO1001327, THYRO1001523, THYRO1001702, THYRO1001725, Y79AA1000426, Y79AA1000521,  
 Y79AA1000959, Y79AA1001013, Y79AA1001264, Y79AA1001328, Y79AA1001427, Y79AA1001430,  
 50 Y79AA1001795, Y79AA1001803, Y79AA1002022,

[0241] Clones whose deduced amino acid sequences were predicted to have transmembrane regions by SOSUI  
 search are as follows: BNGH41000091, HEMBA1000121, HEMBA1000349, HEMBA1000477, HEMBA1000713,  
 HEMBA1000940, HEMBA1000962, HEMBA1001221, HEMBA1001228, HEMBA1001621, HEMBA1002167,  
 HEMBA1002195, HEMBA1002227, HEMBA1002421, HEMBA1003101, HEMBA1003392, HEMBA1003530,  
 55 HEMBA1003732, HEMBA1003945, HEMBA1004391, HEMBA1004454, HEMBA1004797, HEMBA1004982,  
 HEMBA1005449, HEMBA1005522, HEMBA1005545, HEMBA1005698, HEMBA1006171, HEMBA1006299,  
 HEMBA1006311, HEMBA1006335, HEMBA1006357, HEMBA1006430, HEMBA1006724, HEMBA1006960,  
 HEMBB1000407, HEMBB1000447, HEMBB1000567, HEMBB1000679, HEMBB1000905, HEMBB1001026,

HEMBB1001407, HEMBB1001573, HEMBB1001978, HEMBB1002041, HEMBB1002162, HEMBB1002245,  
 HEMBB1002427, HEMBB1002693, MAMMA1000102, MAMMA1000106, MAMMA1000118, MAMMA1000141,  
 MAMMA1000204, MAMMA1000226, MAMMA1000457, MAMMA1000473, MAMMA1000591, MAMMA1000681,  
 MAMMA1000810, MAMMA1000986, MAMMA1001043, MAMMA1001141, MAMMA1001237, MAMMA1001344,  
 5 MAMMA1001893, MAMMA1001957, MAMMA1001978, MAMMA1002070, MAMMA1002091, MAMMA1002095,  
 MAMMA1002633, NT2RM1000580, NT2RM1000855, NT2RM1000858, NT2RM2000410, NT2RM2000565,  
 NT2RM2001626, NT2RM2001939, NT2RM2001941, NT2RM4000444, NT2RM4000587, NT2RM4000648,  
 NT2RM4000997, NT2RM4001325, NT2RM4001735, NT2RM4001768, NT2RM4002352, NT2RP1000050,  
 NT2RP1000181, NT2RP1000261, NT2RP1000300, NT2RP1000448, NT2RP1000551, NT2RP1000613,  
 10 NT2RP1000981, NT2RP1001563, NT2RP2000479, NT2RP2000533, NT2RP2000649,  
 NT2RP2000663, NT2RP2000694, NT2RP2000818, NT2RP2000903, NT2RP2001200, NT2RP2001276,  
 NT2RP2001495, NT2RP2001915, NT2RP2001956, NT2RP2002188, NT2RP2002232, NT2RP2002527,  
 NT2RP2002533, NT2RP2002721, NT2RP2002824, NT2RP2002942, NT2RP2002976, NT2RP2003042,  
 NT2RP2003390, NT2RP2003469, NT2RP2003593, NT2RP2003655, NT2RP2003664, NT2RP2003950,  
 15 NT2RP2004179, NT2RP2004205, NT2RP2004495, NT2RP2004524, NT2RP2004556, NT2RP2004606,  
 NT2RP2004648, NT2RP2004794, NT2RP2005163, NT2RP2005181, NT2RP2005463, NT2RP2005597,  
 NT2RP2005666, NT2RP2005883, NT2RP2005994, NT2RP2006004, NT2RP2006269, NT2RP2006512,  
 NT2RP2006580, NT2RP3000169, NT2RP3000171, NT2RP3000304, NT2RP3000460, NT2RP3000616,  
 NT2RP3000721, NT2RP3000818, NT2RP3000907, NT2RP3000921, NT2RP3001159, NT2RP3001195,  
 20 NT2RP3001240, NT2RP3001271, NT2RP3001322, NT2RP3001388, NT2RP3001560, NT2RP3001592,  
 NT2RP3001650, NT2RP3001738, NT2RP3002015, NT2RP3002311, NT2RP3002342, NT2RP3002411,  
 NT2RP3002790, NT2RP3002836, NT2RP3002900, NT2RP3002958, NT2RP3003000, NT2RP3003354,  
 NT2FP3003532, NT2RP3003535, NT2RP3003614, NT2RP3004025, NT2RP3004075, NT2RP3004083,  
 NT2RP3004090, NT2RP3004130, NT2RP3004294, NT2RP3004309, NT2RP3004345, NT2RP3004406,  
 25 NT2RP3004481, NT2RP3004552, NT2RP4001001, NT2RP4001009, NT2RP4001467, NT2RP4001879,  
 NT2RP4002187, NT2RP4002451, NT2RP4002750, OVARC1000003, OVARC1000105, OVARC1000307,  
 OVARC1000439, OVARC1000553, OVARC1001030, OVARC1001336,  
 OVARC1001570, PLACE1000231, PLACE1000560, PLACE1000740, PLACE1000912, PLACE1000914,  
 PLACE1000927, PLACE1001016, PLACE1001183, PLACE1001231, PLACE1001401, PLACE1001407,  
 30 PLACE1001464, PLACE1001536, PLACE1001564, PLACE1001655, PLACE1001836, PLACE1001918,  
 PLACE1001949, PLACE1002518, PLACE1002726, PLACE1002967, PLACE1003573, PLACE1003737,  
 PLACE1003839, PLACE1003845, PLACE1003852, PLACE1004279, PLACE1004282, PLACE1004441,  
 PLACE1004637, PLACE1004648, PLACE1004816, PLACE1004887, PLACE1005003, PLACE1005005,  
 PLACE1005410, PLACE1005544, PLACE1005569, PLACE1005660, PLACE1005725, PLACE1005745,  
 35 PLACE1005927, PLACE1006290, PLACE1006443, PLACE1006959, PLACE1007096, PLACE1007296,  
 PLACE1007626, PLACE1007881, PLACE1008359, PLACE1008469, PLACE1008716, PLACE1008985,  
 PLACE1009196, PLACE1009279, PLACE1009527, PLACE1009546, PLACE1009600, PLACE1010011,  
 PLACE1010078, PLACE1010445, PLACE1010713, PLACE1010784, PLACE1010968, PLACE1011236,  
 PLACE1011516, PLACE3000181, THYRO1000400, THYRO1000678, THYRO1000776, THYRO1000956,  
 40 THYRO1001102, THYRO1001113, THYRO1001205, THYRO1001237, THYRO1001242, THYRO1001266,  
 THYRO1001327, THYRO1001478, THYRO1001523, THYRO1001641, THYRO1001702, THYRO1001725,  
 Y79AA1000207, Y79AA1000226, Y79AA1000270, Y79AA1000521, Y79AA1000888, Y79AA1001013,  
 Y79AA1001212, Y79AA1001264, Y79AA1001328, Y79AA1001426, Y79AA1001427, Y79AA1001727,  
 Y79AA1001787, Y79AA1001795, Y79AA1001803, Y79AA1002058,  
 45 Y79AA1002129, Y79AA1002213, Y79AA1002373,

**[0242]** Names of clones whose deduced amino acid sequences were predicted to have functional domains by Pfam search, and names of the matched functional domains are shown below. When multiple functional domains matched a clone, each domain name was indicated, separated by a double-slash mark,/.

50 HEMBA1000006//Src homology domain 3  
 HEMBA1000128//SCP-like extracellular Proteins  
 HEMBA1000349//ABC transporters  
 HEMBA1000462//RNA recognition motif. (aka RRM, RBD, or RNP domain)  
 HEMBA1000590//EGF-like domain//von Willebrand factor type A domain  
 55 HEMBA1000671//Zinc finger, C2H2 type  
 HEMBA1000732//EGF-like domain  
 HEMBA1000940//Connexin  
 HEMBA1001221//EGF-like domain//Kazal-type serine protease inhibitor domain

HEMBA1001621//7 transmembrane receptor (rhodopsin family)  
 HEMBA1001878//WD domain, G-beta repeats  
 HEMBA1002048//Zinc finger, C2H2 type  
 HEMBA1002167//Carboxylesterases  
 5 HEMBA1002551//WD domain, G-beta repeats  
 HEMBA1002992//Ubiquitin family  
 HEMBA1003047//CUB domain  
 HEMBA1003120//Zinc finger, C2H2 type  
 HEMBA1003230//EGF-like domain  
 10 HEMBA1003392//Low-density lipoprotein receptor domain class A  
 HEMBA1003497//Zinc finger, C2H2 type  
 HEMBA1004250//Cadherin  
 HEMBA1004391//Fibronectin type III domain//IG superfamily  
 HEMBA1004454//4 transmembrane segments integral membrane proteins  
 15 HEMBA1004785!/'chromo' (CHRromatin Organization MOdifier) domain  
 HEMBA1005246//Zinc finger, C2H2 type  
 HEMBA1005267//Ank repeat  
 HEMBA1005545//7 transmembrane receptor (rhodopsin family)  
 HEMBA1005929//Eukaryotic protein kinase domain  
 20 HEMBA1005945//Mitochondrial carrier proteins  
 HEMBA1006572//Zinc finger, C2H2 type  
 HEMBA1006707//EGF-like domain//von Willebrand factor type A domain  
 HEMBA1006749//EGF-like domain//von Willebrand factor type A domain  
 HEMBA1006770//RNA recognition motif. (aka RRM, RBD, or RNP domain)  
 25 HEMBA1006902//EGF-like domain//von Willebrand factor type A domain  
 HEMBB1000106//Zinc finger, CCHC class  
 HEMBB1000668//WD domain, G-beta repeats  
 HEMBB1000881//Thrombospondin type 1 domain  
 HEMBB1000905//WD domain, G-beta repeats  
 30 HEMBB1002041//EGF-like domain//Kazal-type serine protease inhibitor domain  
 HEMBB1002245//IG superfamily  
 HEMBB1002302//Zinc finger, CCHC class  
 HEMBB1002465//Acyl-CoA dehydrogenases  
 HEMBB1002661//Helix-loop-helix DNA-binding domain  
 35 MAMMA1000204//C2 domain  
 MAMMA1000457//FAD/NAD-binding domain in oxidoreductases  
 MAMMA1000681//7 transmembrane receptor (rhodopsin family)  
 MAMMA1000881//Eukaryotic protein kinase domain//Protein kinase C terminal domain  
 MAMMA1001150//Phorbol esters / diacylglycerol binding domain//Eukaryotic protein kinase domain  
 40 MAMMA1001310//WD domain, G-beta repeats  
 MAMMA1001532//Zinc finger, C2H2 type  
 MAMMA1001615//Helix-loop-helix DNA-binding domain  
 MAMMA1002070//Kringle domain  
 MAMMA1002080//Ras family (contains ATP/GTP binding P-loop)  
 45 MAMMA1002095//E1-E2 ATPases  
 MAMMA1002165//Insulin-like growth factor binding proteins  
 NT2RM1000789//HMG (high mobility group) box  
 NT2RM1000855//eubacterial secY protein  
 NT2RM1000899//Mitochondrial carrier proteins  
 50 NT2RM2000589//PH (pleckstrin homology) domain  
 NT2RM2000632//Helicases conserved C-terminal domain  
 NT2RM2001792//Fibrinogen beta and gamma chains, C-terminal globular domain  
 NT2RM2001902//Eukaryotic protein kinase domain  
 NT2RM2001939//7 transmembrane receptor (rhodopsin family)  
 55 NT2RM2001941//7 transmembrane receptor (rhodopsin family)  
 NT2RM4000284//Class I Histocompatibility antigen, domains alpha 1 and 2  
 NT2RM4000326//Zinc finger, C2H2 type  
 NT2RM4000417//C2 domain

Y79AA1001427//FAD/NAD-binding domain in oxidoreductases  
 Y79AA1001523//Bromodomain//Zinc finger, C3HC4 type (RING finger)  
 Y79AA1001530//Tubulin  
 Y79AA1001727//IG superfamily  
 5 Y79AA1001787//E1-E2 ATPases  
 Y79AA1001799//Mitochondrial carrier proteins  
 Y79AA1002022//IG superfamily  
 Y79AA1002381//Eukaryotic protein kinase domain

## 10 EXAMPLE 17

Functional categories based on the full-length nucleotide sequences

15 **[0243]** Prediction of functions of proteins encoded by the clones and the categorization thereof were performed based on the results of homology search (see homology search result 10) of the databases, GenBank, Swiss-Prot and UniGene for the full-length nucleotide sequences of 826 clones as well as based on the results of domain search (see Example 16) of the deduced amino acid sequences encoded by the full-length nucleotide sequences. (HEMBA1005337, NT2RM1000407, NT2RM2001767, and NT2RP3003939 were excluded because of the absence of full-length sequence.)

20 **[0244]** There are 611 clones that presumably encode proteins belonging to any of categories of secretory and/or membrane proteins, glycoprotein-associated proteins, signal transduction-associated proteins, transcription-associated proteins and disease-associated proteins.

25 **[0245]** The clones presumably encoding proteins categorized into secretory and/or membrane proteins are those which matched the full-length sequences of Swiss-Prot database with keywords "growth factor", "cytokine", "hormone", "signal", "transmembrane", "membrane", "extracellular matrix", "receptor", "G-protein coupled receptor", "ionic channel", "voltage-gated channel", "calcium channel", "cell adhesion", "collagen" or "connective tissue"; those which matched the data, suggesting that the proteins are secretory and/or membrane proteins; or those which matched the full-length sequences of GenBank or UniGene database with similar description; and, further, those predicted to have an N-terminal signal sequence or a transmembrane region as a result of domain search for the amino acid sequences deduced from the full-length nucleotide sequences.

30 **[0246]** The clones presumably encoding proteins categorized into glycoprotein-associated proteins are those which matched the full-length sequences of Swiss-Prot database with the keywords "glycoprotein"; those which matched the data, suggesting that the proteins are glycoprotein; or those which matched the full-length sequences of GenBank or UniGene database.

35 **[0247]** The clones presumably encoding proteins categorized into signal transduction-associated proteins are those which matched the full-length sequences of Swiss-Prot database with the keywords "serine/threonine-protein kinase", "tyrosine-protein kinase" or "SH3 domain"; those which matched the data, suggesting that the proteins are signal transduction-associated proteins (for example, "ADP-ribosylation factor"); or those which matched the full-length sequences of GenBank or UniGene database with similar description.

40 **[0248]** The clones presumably encoding proteins categorized into transcription-associated proteins are those which matched the full-length sequences of Swiss-Prot database with the keywords "transcription regulation", "zinc finger" or "homeobox"; those which matched the data, suggesting that the proteins are transcription-associated proteins; or those which matched the full-length sequences of GenBank or UniGene database with similar description.

45 **[0249]** The clones presumably encoding proteins categorized into disease-associated proteins are those which matched the full-length sequences of Swiss-Prot database with the keywords "disease mutation" or "syndrome"; those which matched the data, suggesting that the proteins are disease-associated proteins; or those which matched the full-length sequences of Swiss-Prot database and GenBank or UniGene database where the matched sequences are those of genes or proteins which had been deposited in the database of Online Mendelian Inheritance in Man (OMIM) (<http://www.ncbi.nlm.nih.gov/Omim/>), which is a database of human genes and diseases.

50 **[0250]** The following 437 clones were categorized into secretory and/or membrane proteins.

BNGH41000020, BNGH41000087, BNGH41000091, HEMBA1000121, HEMBA1000128, HEMBA1000349,  
 HEMBA1000477, HEMBA1000590, HEMBA1000713, HEMBA1000732, HEMBA1000745, HEMBA1000835,  
 HEMBA1000940, HEMBA1000962, HEMBA1001221, HEMBA1001228, HEMBA1001621, HEMBA1002131,  
 HEMBA1002163, HEMBA1002167, HEMBA1002178, HEMBA1002195, HEMBA1002227, HEMBA1002420,  
 55 HEMBA1002421, HEMBA1002767, HEMBA1003047, HEMBA1003101, HEMBA1003230, HEMBA1003392,  
 HEMBA1003530, HEMBA1003602, HEMBA1003732, HEMBA1003945, HEMBA1004110, HEMBA1004250,  
 HEMBA1004391, HEMBA1004444, HEMBA1004454, HEMBA1004505, HEMBA1004797, HEMBA1004982,  
 HEMBA1005070, HEMBA1005449, HEMBA1005522, HEMBA1005545, HEMBA1005698, HEMBA1005945,



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	HEMBA1006171,	HEMBA1006299,	HEMBA1006311,	HEMBA1006335,	HEMBA1006357,	HEMBA1006430,
	HEMBA1006482,	HEMBA1006707,	HEMBA1006724,	HEMBA1006749,	HEMBA1006902,	HEMBA1006960,
	HEMBA1007241,	HEMBB1000407,	HEMBB1000447,	HEMBB1000567,	HEMBB1000679,	HEMBB1000881,
	HEMBB1001026,	HEMBB1001048,	HEMBB1001407,	HEMBB1001530,	HEMBB1001573,	HEMBB1001847,
5	HEMBB1001978,	HEMBB1002041,	HEMBB1002162,	HEMBB1002245,	HEMBB1002427,	HEMBB1002693,
	MAMMA1000102,	MAMMA1000106,	MAMMA1000118,	MAMMA1000141,	MAMMA1000204,	MAMMA1000226,
	MAMMA1000457,	MAMMA1000473,	MAMMA1000496,	MAMMA1000591,	MAMMA1000681,	MAMMA1000810,
	MAMMA1000986,	MAMMA1000994,	MAMMA1001043,	MAMMA1001141,	MAMMA1001237,	MAMMA1001344,
	MAMMA1001418,	MAMMA1001893,	MAMMA1001957,	MAMMA1001978,		
10	MAMMA1002070,	MAMMA1002091,	MAMMA1002095,	MAMMA1002165,	MAMMA1002234,	MAMMA1002586,
	MAMMA1002633,	MAMMA1003126,	NT2RM1000462,	NT2RM1000542,	NT2RM1000580,	NT2RM1000855,
	NT2RM1000858,	NT2RM1000899,	NT2RM2000241,	NT2RM2000410,	NT2RM2000423,	NT2RM2000565,
	NT2RM2001626,	NT2RM2001792,	NT2RM2001939,	NT2RM2001941,	NT2RM4000198,	NT2RM4000284,
	NT2RM4000417,	NT2RM4000444,	NT2RM4000587,	NT2RM4000593,	NT2RM4000648,	NT2RM4000761,
15	NT2RM4000997,	NT2RM4001325,	NT2RM4001735,	NT2RM4001768,	NT2RM4001843,	NT2RM4002352,
	NT2RP1000050,	NT2RP1000181,	NT2RP1000261,	NT2RP1000300,	NT2RP1000325,	NT2RP1000448,
	NT2RP1000551,	NT2RP1000613,	NT2RP1000981,	NT2RP1001004,	NT2RP1001563,	NT2RP2000479,
	NT2RP2000533,	NT2RP2000616,	NT2RP2000649,	NT2RP2000663,	NT2RP2000694,	NT2RP2000818,
	NT2RP2000903,	NT2RP2001200,	NT2RP2001276,	NT2RP2001480,	NT2RP2001495,	NT2RP2001514,
20	NT2RP2001755,	NT2RP2001915,	NT2RP2001956,	NT2RP2002063,	NT2RP2002188,	NT2RP2002232,
	NT2RP2002527,	NT2RP2002533,	NT2RP2002721,	NT2RP2002824,	NT2RP2002942,	NT2RP2002976,
	NT2RP2003042,	NT2RP2003210,	NT2RP2003383,	NT2RP2003390,	NT2RP2003469,	NT2RP2003593,
	NT2RP2003655,	NT2RP2003664,	NT2RP2003950,	NT2RP2004179,	NT2RP2004205,	NT2RP2004495,
	NT2RP2004524,	NT2RP2004556,	NT2RP2004606,	NT2RP2004648,	NT2RP2004794,	NT2RP2005027,
25	NT2RP2005163,	NT2RP2005181,	NT2RP2005378,	NT2RP2005463,	NT2RP2005541,	NT2RP2005597,
	NT2RP2005666,	NT2RP2005883,	NT2RP2005994,	NT2RP2006004,		
	NT2RP2006042,	NT2RP2006269,	NT2RP2006512,	NT2RP2006580,	NT2RP3000169,	NT2RP3000171,
	NT2RP3000304,	NT2RP3000436,	NT2RP3000460,	NT2RP3000616,	NT2RP3000676,	NT2RP3000721,
	NT2RP3000818,	NT2RP3000907,	NT2RP3000921,	NT2RP3001012,	NT2RP3001159,	NT2RP3001195,
30	NT2RP3001240,	NT2RP3001271,	NT2RP3001322,	NT2RP3001388,	NT2RP3001560,	NT2RP3001592,
	NT2RP3001650,	NT2RP3001738,	NT2RP3001858,	NT2RP3002015,	NT2RP3002160,	NT2RP3002311,
	NT2RP3002342,	NT2RP3002411,	NT2RP3002737,	NT2RP3002790,	NT2RP3002836,	NT2RP3002900,
	NT2RP3002958,	NT2RP3003000,	NT2RP3003076,	NT2RP3003354,	NT2RP3003532,	NT2RP3003535,
	NT2RP3003614,	NT2RP3004025,	NT2RP3004075,	NT2RP3004083,	NT2RP3004130,	NT2RP3004133,
35	NT2RP3004309,	NT2RP3004345,	NT2RP3004406,	NT2RP3004481,	NT2RP3004552,	NT2RP3004625,
	NT2RP3004647,	NT2RP4001001,	NT2RP4001009,	NT2RP4001467,	NT2RP4001879,	NT2RP4002187,
	NT2RP4002451,	NT2RP4002750,	OVARC1000003,	OVARC1000105,	OVARC1000298,	OVARC1000307,
	OVARC1000313,	OVARC1000410,	OVARC1000439,	OVARC1000553,	OVARC1000811,	OVARC1000873,
	OVARC1000956,	OVARC1001030,	OVARC1001163,	OVARC1001336,	OVARC1001570,	OVARC1001607,
40	OVARC1001725,	OVARC1001991,	PLACE1000033,	PLACE1000231,	PLACE1000560,	PLACE1000740,
	PLACE1000912,	PLACE1000914,	PLACE1000927,	PLACE1001016,	PLACE1001123,	PLACE1001183,
	PLACE1001231,	PLACE1001340,	PLACE1001401,	PLACE1001407,	PLACE1001464,	PLACE1001516,
	PLACE1001536,	PLACE1001564,	PLACE1001655,	PLACE1001795,		
	PLACE1001836,	PLACE1001918,	PLACE1001949,	PLACE1002080,	PLACE1002095,	PLACE1002355,
45	PLACE1002374,	PLACE1002518,	PLACE1002547,	PLACE1002726,	PLACE1002905,	PLACE1002911,
	PLACE1002967,	PLACE1003407,	PLACE1003573,	PLACE1003737,	PLACE1003772,	PLACE1003839,
	PLACE1003845,	PLACE1003852,	PLACE1004279,	PLACE1004282,	PLACE1004441,	PLACE1004450,
	PLACE1004482,	PLACE1004520,	PLACE1004630,	PLACE1004637,	PLACE1004648,	PLACE1004816,
	PLACE1005003,	PLACE1005005,	PLACE1005031,	PLACE1005383,	PLACE1005410,	PLACE1005426,
50	PLACE1005544,	PLACE1005569,	PLACE1005660,	PLACE1005725,	PLACE1005745,	PLACE1005878,
	PLACE1005927,	PLACE1006071,	PLACE1006093,	PLACE1006208,	PLACE1006277,	PLACE1006290,
	PLACE1006443,	PLACE1006716,	PLACE1006809,	PLACE1006959,	PLACE1007081,	PLACE1007096,
	PLACE1007296,	PLACE1007626,	PLACE1007845,	PLACE1007881,	PLACE1008359,	PLACE1008469,
	PLACE1008716,	PLACE1008744,	PLACE1008985,	PLACE1009067,	PLACE1009196,	PLACE1009279,
55	PLACE1009527,	PLACE1009546,	PLACE1009600,	PLACE1009982,	PLACE1010011,	PLACE1010078,
	PLACE1010251,	PLACE1010445,	PLACE1010713,	PLACE1010784,	PLACE1010827,	PLACE1010968,
	PLACE1011116,	PLACE1011181,	PLACE1011236,	PLACE1011516,	PLACE1011708,	PLACE3000181,
	PLACE3000213,	PLACE4000354,	SKNMC1000004,	SKNMC1000014,	SKNMC1000082,	THYRO1000036,

- THYRO1000099, THYRO1000196, THYRO1000400, THYRO1000584, THYRO1000678, THYRO1000776,  
 THYRO1000795, THYRO1000956, THYRO1001102, THYRO1001113,  
 THYRO1001205, THYRO1001237, THYRO1001242, THYRO1001266, THYRO1001327, THYRO1001456,  
 THYRO1001478, THYRO1001523, THYRO1001529, THYRO1001641, THYRO1001702, THYRO1001725,  
 5 Y79AA1000207, Y79AA1000226, Y79AA1000270, Y79AA1000426, Y79AA1000521, Y79AA1000876,  
 Y79AA1000888, Y79AA1000959, Y79AA1001013, Y79AA1001212, Y79AA1001264, Y79AA1001328,  
 Y79AA1001426, Y79AA1001427, Y79AA1001430, Y79AA1001727, Y79AA1001787, Y79AA1001795,  
 Y79AA1001799, Y79AA1001803, Y79AA1002022, Y79AA1002058, Y79AA1002129, Y79AA1002213,  
 Y79AA1002373,  
 10 **[0251]** The following 146 clones were categorized into glycoprotein-associated proteins.  
 BNGH41000087, BNGH41000091, HEMBA1000349, HEMBA1000590, HEMBA1000745, HEMBA1000835,  
 HEMBA1001221, HEMBA1001228, HEMBA1001621, HEMBA1002131, HEMBA1002178, HEMBA1002421,  
 HEMBA1002767, HEMBA1003230, HEMBA1003392, HEMBA1004250, HEMBA1004391, HEMBA1004444,  
 HEMBA1004505, HEMBA1005449, HEMBA1005522, HEMBA1005545, HEMBA1006707, HEMBA1006749,  
 15 HEMBA1006902, HEMBB1000679, HEMBB1000881, HEMBB1001048, HEMBB1002120, HEMBB1002245,  
 HEMBB1002427, MAMMA1000102, MAMMA1000591, MAMMA1000681, MAMMA1001043, MAMMA1001237,  
 MAMMA1002070, MAMMA1002586, MAMMA1003126, NT2RM1000462, NT2RM1000580, NT2RM2001792,  
 NT2RM2001818, NT2RM2001939, NT2RM2001941, NT2RM4000198, NT2RM4000284, NT2RM4000417,  
 NT2RM4000648, NT2RM4000997, NT2RM4001325, NT2RM4002352, NT2RP1000613, NT2RP1000981,  
 20 NT2RP1001004, NT2RP2000616, NT2RP2000694, NT2RP2000903, NT2RP2001480, NT2RP2001755,  
 NT2RP2002533, NT2RP2003042, NT2RP2003210, NT2RP2004205, NT2RP2004606, NT2RP2005027,  
 NT2RP2005181, NT2RP2005541, NT2RP2005597, NT2RP2005883, NT2RP2006004, NT2RP2006042,  
 NT2RP2006269, NT2RP3000304, NT2RP3000616, NT2RP3000921, NT2RP3001650, NT2RP3002160,  
 NT2RP3002737, NT2RP3002958, NT2RP3003000, NT2RP3003532, NT2RP3004130, NT2RP3004133,  
 25 NT2RP3004481, NT2RP3004552, NT2RP3004640, NT2RP4000108, NT2RP4001467, NT2RP4002750,  
 OVARC1000003, OVARC1000553, OVARC1000811, OVARC1000873, OVARC1001336, OVARC1001607,  
 OVARC1001991, PLACE1000033, PLACE1000740, PLACE1001016,  
 PLACE1001123, PLACE1001231, PLACE1001464, PLACE1001655, PLACE1001836, PLACE1002355,  
 PLACE1002374, PLACE1002905, PLACE1002911, PLACE1003573, PLACE1003737, PLACE1003772,  
 30 PLACE1003839, PLACE1004282, PLACE1004441, PLACE1004450, PLACE1004520, PLACE1004648,  
 PLACE1005003, PLACE1005426, PLACE1006071, PLACE1006073, PLACE1006290, PLACE1007081,  
 PLACE1007845, PLACE1008716, PLACE1008744, PLACE1008985, PLACE1010251, PLACE1010784,  
 PLACE1010968, PLACE1011116, PLACE3000181, PLACE3000213, PLACE4000354, THYRO1000036,  
 THYRO1000196, THYRO1000584, THYRO1000956, THYRO1001266, Y79AA1000270, Y79AA1000426,  
 35 Y79AA1001727, Y79AA1001795, Y79AA1002022, Y79AA1002213,  
**[0252]** The following 55 clones were categorized into signal transduction-associated proteins.  
 HEMBA1000006, HEMBA1002195, HEMBA1002227, HEMBA1002551, HEMBA1005084, HEMBA1005929,  
 HEMBA1006658, HEMBA1006916, MAMMA1000881, MAMMA1001150, MAMMA1001310, MAMMA1002142,  
 NT2RM2001902, NT2RP1001020, NT2RP1001031, NT2RP2001469, NT2RP2001529, NT2RP2001769,  
 40 NT2RP2003179, NT2RP2003545, NT2RP2004670, NT2RP3000011, NT2RP3000022, NT2RP3000172,  
 NT2RP3000201, NT2RP3000820, NT2RP3003527, NT2RP3003849, NT2RP3003874, NT2RP3004067,  
 NT2RP4000634, NT2RP4000962, OVARC1000255, OVARC1000529, OVARC1000916, OVARC1001338,  
 OVARC1001569, PLACE1002329, PLACE1003135, PLACE1003598, PLACE1005519, PLACE1006208,  
 PLACE1008282, PLACE1008297, PLACE1010081, PLACE1011364, PLACE1011824, THYRO1001457,  
 45 THYRO1001593, THYRO1001700, THYRO1001770, Y79AA1000777, Y79AA1000967, Y79AA1002376,  
 Y79AA1002381,  
**[0253]** The following 80 clones were categorized into transcription-associated proteins.  
 HEMBA1000462, HEMBA1000671, HEMBA1001297, HEMBA1001390, HEMBA1001886, HEMBA1002048,  
 HEMBA1003120, HEMBA1003497, HEMBA1004785, HEMBA1005230, HEMBA1005246, HEMBA1006276,  
 50 HEMBA1006572, HEMBA1007226, HEMBB1000106, HEMBB1000905, HEMBB1001959, HEMBB1002051,  
 HEMBB1002661, MAMMA1001094, MAMMA1001532, MAMMA1001615, NT2RM1000789, NT2RM2000632,  
 NT2RM2000773, NT2RM4000326, NT2RP1000271, NT2RP1000468, NT2RP2000092, NT2RP2000610,  
 NT2RP2000712, NT2RP2000739, NT2RP2001538, NT2RP2001662, NT2RP2001817, NT2RP2001948,  
 NT2RP2002564, NT2RP2002974, NT2RP2003138, NT2RP2003302, NT2RP2003940, NT2RP2004108,  
 55 NT2RP2004847, NT2RP2005247, NT2RP2005391, NT2RP2005535, NT2RP2005774, NT2RP2005941,  
 NT2RP2006092, NT2RP3000148, NT2RP3000232, NT2RP3000378, NT2RP3000652, NT2RP3001976,  
 NT2RP3004090, NT2RP3004119, NT2RP3004294, OVARC1001049, OVARC1001086, OVARC1001132,  
 OVARC1001807, PLACE1000258, PLACE1000442, PLACE1000907, PLACE1003529, PLACE1004166,

PLACE1004168, PLACE1004887, PLACE1005250, PLACE1005682, PLACE1006079, PLACE1008549,  
PLACE1011407, PLACE1011978, THYRO1000580, Y79AA1000030, Y79AA1001090, Y79AA1001523,  
Y79AA1002334, Y79AA1002378,

[0254] The following 85 clones were categorized into disease-associated proteins.

5 BNGH41000020, HEMBA1000349, HEMBA1000590, HEMBA1000671, HEMBA1000835, HEMBA1001184,  
HEMBA1001228, HEMBA1001886, HEMBA1003120, HEMBA1004250, HEMBA1005246, HEMBA1005267,  
HEMBA1006707, HEMBA1006749, HEMBA1006902, HEMBA1006916, HEMBA1007013, HEMBB1002120,  
MAMMA1000204, MAMMA1002080, NT2RM2000632, NT2RM2001126, NT2RM2001558, NT2RP1000271,  
NT2RP1000465, NT2RP1000579, NT2RP2000447, NT2RP2000514, NT2RP2000739, NT2RP2001223,  
10 NT2RP2001529, NT2RP2001562, NT2RP2002674, NT2RP2003369, NT2RP2004108, NT2RP2004205,  
NT2RP2005535, NT2RP2005941, NT2RP2006004, NT2RP3000059, NT2RP3000125, NT2RP3000201,  
NT2RP3000232, NT2RP3000616, NT2RP3000677, NT2RP3000838, NT2RP3000921, NT2RP3001542,  
NT2RP3002286, NT2RP3002721, NT2RP3002737, NT2RP3002738, NT2RP3004481, OVARC1000208,  
OVARC1000275, OVARC1000331, OVARC1000410, OVARC1001086, OVARC1001132, OVARC1001607,  
15 OVARC1001725, OVARC1001952, PLACE1000258, PLACE1000442, PLACE1000907, PLACE1001100,  
PLACE1001500, PLACE1002905, PLACE1002967, PLACE1003407, PLACE1003428, PLACE1005005,  
PLACE1005239, PLACE1005815, PLACE1007028, PLACE1008716, PLACE1011407, PLACE1011978,  
PLACE2000118, THYRO1000580, THYRO1000866, THYRO1001071, THYRO1001478, Y79AA1001062,  
Y79AA1001530,

20 [0255] Out of them, the following 67 clones are those which matched the data of Swiss-Prot database and GenBank or UniGene database, genes or proteins which had been deposited in the database of Online Mendelian Inheritance in Man (OMIM) (<http://www.ncbi.nlm.nih.gov/Omim/>), which is a database of human genes and diseases. (The corresponding OMIM numbers are indicated after the clone names.)

HEMBA1000349(\*600046), HEMBA1000590(\*603897), HEMBA1000671(\*602277), HEMBA1001886(\*603899),  
25 HEMBA1003120(\*602277), HEMBA1004250(\*600976), HEMBA1005246(\*602291), HEMBA1005267(\*106410),  
HEMBA1006707(\*603897), HEMBA1006749(\*603897), HEMBA1006902(\*603897), HEMBA1006916(\*601524),  
HEMBA1007013(\*603730), HEMBB1002120(\*603367), MAMMA1002080(\*602672), NT2RM2001126(\*603785),  
NT2RM2001558(\*604689), NT2RP1000271(\*603899), NT2RP1000465(\*602231), NT2RP2000447(\*602580),  
NT2RP2000514(\*602431), NT2RP2000739(\*194558), NT2RP2001223(\*603558), NT2RP2001529(\*603289),  
30 NT2RP2001562(\*603371), NT2RP2002674(\*132811), NT2RP2003369(\*179555), NT2RP2004108(\*601260),  
NT2RP2004205(\*601610), NT2RP2005535(\*603899), NT2RP2006004(\*600245), NT2RP3000059(\*106410),  
NT2RP3000125(\*180202), NT2RP3000201(\*604666), NT2RP3000232(\*602277), NT2RP3000616(\*600245),  
NT2RP3000677(\*142765), NT2RP3000838(\*190370), NT2RP3001542(\*191161), NT2RP3002286(\*604331),  
NT2RP3002721(\*118950), NT2RP3002738(\*602265), NT2RP3004481(\*601610), OVARC1000208(\*603603),  
35 OVARC1000275(\*125647), OVARC1000331(\*139265), OVARC1000410(\*603874), OVARC1001086(\*603862),  
OVARC1001725(\*603046), OVARC1001952(\*190370), PLACE1000258(\*603971), PLACE1000442(\*601260),  
PLACE1000907(\*194558), PLACE1001500(\*603781), PLACE1002905(\*125950), PLACE1003428(\*603570),  
PLACE1005005(\*603124), PLACE1005239(\*603365), PLACE1007028(\*602131), PLACE1011407(\*602277),  
PLACE1011978(\*603971), PLACE2000118(\*301000), THYRO1000580(\*602277), THYRO1000866(\*604045),  
40 THYRO1001071(\*603533), Y79AA1001062(\*191161), Y79AA1001530(\*602662),

[0256] Out of 215 clones excluding the above-mentioned clones, HEMBB1000668 and NT2RM4001377 presumably belong to a group of signal transduction-associated proteins, based on the results of domain search by Pfam.

[0257] HEMBB1002302 presumably belong to a group of transcription-associated proteins, based on the results of domain search by Pfam.

45 [0258] In the 437 clones categorized into secretory and/or transmembrane proteins on the basis of their full-length sequences, 410 clones were also predicted to encode proteins having functions of secretory and/or membrane proteins on the basis of their partial nucleotide sequences (5' sequences). In the 146 clones categorized into glycoprotein-associated proteins on the basis of their full-length sequences, 124 clones were also predicted to encode proteins having functions of glycoprotein-associated proteins on the basis of their partial nucleotide sequences. In the 57 clones  
50 categorized into signal transduction-associated proteins on the basis of their full-length sequences, 46 clones were also predicted to encode proteins having functions of signal transduction-associated proteins on the basis of their partial nucleotide sequences. In the 81 clones categorized into transcription-associated proteins on the basis of their full-length sequences, 57 clones were also predicted to encode proteins having functions of transcription-associated proteins on the basis of their partial nucleotide sequences. In the 85 clones categorized into disease-associated proteins  
55 on the basis of their full-length sequences, 6 clones were also predicted to encode proteins having functions of disease-associated proteins on the basis of their partial nucleotide sequences. The number of clones which were predicted to encode disease-associated proteins based on the full-length nucleotide sequences is much greater than that predicted based on the partial sequences. The reason is that the full-length sequences were categorized by using the data found

in the OMIM database into the category of disease-associated proteins.

**[0259]** When the predicted functions based on the partial sequences were different from those based on the full-length sequences, several reasons were presumed; the ORF is too short in the partial sequence as compared with that of the full-length sequence; alternatively, P value for the partial sequence was greater than that for the full-length, that is, as compared with the probability of occurrence of the predicted function found in the full-length sequence, the probability was lower in the partial sequence. A protein does not always belong solely to a single category of the above-described functional categories, and therefore, additional functions can be found for the cDNA clones by further analyses.

**[0260]** It is unclear, by the analyses for the full-length sequences so far, whether or not the remaining 212 clones encode proteins belonging to any of the categories of secretory and/or membrane proteins, glycoprotein-associated proteins, signal transduction-associated proteins, transcription-associated proteins or disease-associated proteins. Nonetheless, the functions which were predicted based on the partial sequences can be verified by further analyses.

**[0261]** Among the 212 clones, there are 38 clones that presumably belong to the category of enzymes and/or metabolism-associated proteins, cell division- and/or cell proliferation-associated proteins, cytoskeleton-associated proteins, nuclear proteins, DNA-and/or RNA-binding proteins, ASP- and/or GTP-binding proteins, protein synthesis- and/or protein transport-associated proteins, or cellular defense-associated proteins. The clones containing results of homology search of Swiss-Prot database were categorized by considering the keywords and mentioned items in the matching data. The clones containing results of homology search of GenBank or UniGene database were categorized by considering the definitions and mentioned items in the matching data.

When the matching data contained keywords such as "metabolism", "oxidoreductase" and "E.C. No. (Enzyme commission number)", the clones were herein defined as clones presumably belonging to the category of enzymes and/or metabolism-associated proteins. When the matching data contained keywords such as "cell division", "cell cycle", "mitosis", "chromosomal protein", "cell growth" and "apoptosis", the clones were herein defined as clones presumably belonging to the category of cell division- or cell proliferation-associated proteins. When the matching data contained keywords such as "structural protein", "cytoskeleton", "actin-binding" and "microtubules", the clones were herein defined as clones presumably belonging to the category of cytoskeleton-associated proteins. When the matching data contained keywords such as "nuclear protein", the clones were herein defined as clones presumably belonging to the category of nuclear proteins. When the matching data contained keywords such as "DNA-binding" and "RNA-binding", the clones were herein defined as clones presumably belonging to the category of DNA- or RNA-binding proteins. When the matching data contained keywords such as "ATP-binding" and "GTP-binding", the clones were herein defined as clones presumably belonging to the category of ATP- and/or GTP-binding proteins. When the matching data contained keywords such as "translation regulation", "protein biosynthesis", "amino-acid biosynthesis", "ribosomal protein", "protein transport" and "signal recognition particle", the clones were herein defined as clones presumably belonging to the category of protein synthesis- and/or protein transport-associated proteins. When the matching data contained keywords such as "heat shock", "DNA repair" and "DNA damage", the clones were herein defined as clones presumably belonging to the category of cellular defense-associated proteins.

**[0262]** The following 10 clones presumably belong to enzymes and/or metabolism-associated proteins.

HEMBA1003315, HEMBB1002465, MAMMA1000614, NT2RP2000178, NT2RP2001388, NT2RP2001903, NT2RP2002304, NT2RP2005878, NT2RP3001685, PLACE1006219

**[0263]** The following 4 clones presumably belong to cell division-associated and/or cell proliferation-associated proteins.

MAMMA1000403, NT2RM2000497, NT2RP2000394, Y79AA1002121

**[0264]** The following 6 clones presumably belong to cytoskeleton-associated proteins.

MAMMA1001609, NT2RM2000589, NT2RP3000063, PLACE1004078, PLACE 1004492, PLACE 1008657

**[0265]** The following 7 clones presumably belong to nuclear proteins.

HEMBA1001878, HEMBA1002992, MAMMA1000614, NT2RM4000965, NT2RM2001738, NT2RP2001388, Y79AA1002121

**[0266]** The following 5 clones presumably belong to DNA- and/or RNA-binding proteins.

HEMBA1003072, HEMBA1006770, HEMBA1007332, NT2RM2000497, Y79AA1002121

**[0267]** The following 7 clones presumably belong to ATP- and/or GTP-binding proteins.

HEMBA1002316, MAMMA1001609, NT2RM2000306, NT2RM2000497, NT2RP2000178, NT2RP3003729, PLACE1004305

**[0268]** The following 7 clones presumably belong to protein synthesis- and/or protein transport-associated proteins.

NT2RM4000965, NT2RP2005069, NT2RP3000481, NT2RP3000789, NT2RP4001877, OVARC1001833, OVARC1002058,

**[0269]** The following clone presumably belongs to cellular defense-associated proteins.

PLACE 1005539

**[0270]** Although it is unclear whether or not 26 out of 174 clones other than the above-mentioned clones belong to

any of the above-described categories, these clones are predicted to have some functions, based on the homology search using their full-length sequences. The clone names and the gene definitions found in the result of homology search are shown below, separated by a double-slash, //

- 5 HEMBA1000634//Homo sapiens T-cell activation protein (PGR1) gene, complete cds.  
 HEMBA1002524//Human MHC Class I region proline rich protein mRNA, complete cds.  
 HEMBA1003399//MVP1 PROTEIN,  
 HEMBA1005489//Mus musculus semaphorin cytoplasmic domain-associated protein 3A (Semcap3) mRNA, complete cds.
- 10 HEMBB1000542//Mus musculus bromodomain-containing protein BP75 mRNA, complete cds.  
 MAMMA1000788//Bos taurus P14 (p14) mRNA, complete cds.  
 MAMMA1002128//ABC1 PROTEIN HOMOLOG PRECURSOR.  
 NT2RM2000514//Homo sapiens F-box protein Fbx21 (FBX21) mRNA, complete cds.  
 NT2RM2000622//Mus musculus F-box protein FBL10 mRNA, partial cds.
- 15 NT2RM4000100//Homo Sapiens Leman coiled-coil protein (LCCP) mRNA, complete cds.  
 NT2RP2005425//Homo sapiens mRNA for AKAP450 protein.  
 NT2RP3001170//Mus musculus activity-dependent neuroprotective protein (Adnp) mRNA, complete cds.  
 NT2RP3002571//Bos taurus mRNA for lyncein.  
 NT2RP3004557//Human Ki nuclear autoantigen mRNA, complete cds.
- 20 OVARC1001596//Homo sapiens Arf-like 2 binding protein BART1 mRNA, complete cds.  
 PLACE1002153//Homo sapiens TACC2 protein (TACC2) mRNA, partial cds.  
 PLACE1003163//Homo sapiens DBI-related protein mRNA, complete cds.  
 PLACE1005736//Human mRNA for BAS-GRIP protein.  
 PLACE1007702//Mus musculus TRA1 mRNA, complete cds.
- 25 PLACE1011045//Homo sapiens E1-like protein mRNA, complete cds.  
 THYRO1000061//Mus musculus mRNA for UBE-1c1, UBE-1c2, UBE-1c3, complete cds.  
 THYRO1000964//Drosophila melanogaster Felle associated protein Pellino (Pli) mRNA, complete cds.  
 Y79AA1000776//Mus musculus mRNA for GSG1, complete cds.  
 Y79AA1001056//Homo sapiens MAID protein mRNA, complete cds.
- 30 Y79AA1001272//Homo sapiens retinoic acid repressible protein (RARG-1) mRNA, complete cds.  
 Y79AA1001793//Mus musculus mRNA for GSG1, complete cds.

[0271] So far, useful information for presuming the functions are unavailable for the remaining 148 clones, of which names are listed below.

- 35 HEMBA1000275, HEMBA1000300, HEMBA1000443, HEMBA1000875, HEMBA1000907, HEMBA1001272,  
 HEMBA1001296, HEMBA1001563, HEMBA1002164, HEMEA1002239, HEMBA1002985, HEMBA1003294,  
 HEMBA1003487, HEMBA1004007, HEMBA1004067, HEMBA1004085, HEMBA1004952, HEMBA1004971,  
 HEMBA1005145, HEMBA1005430, HEMBA1005913, HEMBA1006016, HEMBA1006517, HEMBA1006544,  
 HEMBA1006912, HEMBA1007057, HEMBA1007063, HEMBA1007291, HEMBB1000276, HEMBB1000309,  
 40 HEMBB1000642, HEMBB1001200, HEMBB1001547, HEMBB1002039, HEMBB1002228, HEMBB1002663,  
 MAMMA1000046, MAMMA1000449, MAMMA1000528, MAMMA1000652, MAMMA1000706, MAMMA1000814,  
 MAMMA1001066, MAMMA1001284, MAMMA1001623, MAMMA1001634, MAMMA1001901, MAMMA1002087,  
 MAMMA1002205, MAMMA1002224, NT2RM2000582, NT2RM2001643, NT2RM4000115, NT2RM4000295,  
 NT2RM4001321, NT2RP1000002, NT2RP1000239, NT2RP1000679, NT2RP1000740, NT2RP1000903,  
 45 NT2RP2000240, NT2RP2001878, NT2RP2001921, NT2RP2002015, NT2RP2002409, NT2RP2002510,  
 NT2RP2003599, NT2RP2003931, NT2RP2004069, NT2RP2004141, NT2RP2004447, NT2RP2004837,  
 NT2RP2005514, NT2RP2005632, NT2RP2005887, NT2RP2006099, NT2RP2006134, NT2RP3000427,  
 NT2RP3000444, NT2RP3000645, NT2RP3000871, NT2RP3001044, NT2RP3001061, NT2RP3001754,  
 NT2RP3002281, NT2RP3002324, NT2RP3002353, NT2RP3002409, NT2RP3002448, NT2RP3002664,  
 50 NT2RP3002887, NT2RP3002983, NT2RP3003448, NT2RP3003469, NT2RP3003473, NT2RP3003559,  
 NT2RP3003963, NT2RP3004000, NT2RP3004202, NT2RP3004321,  
 NT2RP3004355, NT2RP3004374, NT2RP4002715, OVARC1000090, OVARC1000137, OVARC1000467,  
 OVARC1000775, OVARC1000853, OVARC1000995, OVARC1001222, OVARC1001260, OVARC1001727,  
 OVARC1002178, PLACE1000986, PLACE1001114, PLACE1001229, PLACE1001788, PLACE1003438,  
 55 PLACE1003460, PLACE1003644, PLACE1004028, PLACE1004199, PLACE1004519, PLACE1005601,  
 PLACE1005669, PLACE1005768, PLACE1006515, PLACE1006786, PLACE1007040, PLACE1007077,  
 PLACE1007591, PLACE1007971, PLACE1008984, PLACE1009735, PLACE2000219, PLACE4000455,  
 THYRO1000846, THYRO1000999, THYRO1001063, THYRO1001128, THYRO1001471, THYRO1001495,

THYRO1001608, THYRO1001803, Y79AA1000127, Y79AA1000750, Y79AA1001592, Y79AA1001863,

# EXAMPLE 18

## 5 Expression frequency analysis using PCR

[0272] Many genes acting at the downstream of TNF- and IL-1. among inflammation-associated cytokines have been previously identified. The respective stimulations are transduced through independent pathways of signaling cascade. There exists another signaling cascade for both stimulations, wherein NF- $\kappa$ B is a common transducing molecule shared by the two stimulations (J. Leukoc. Biol., 1994, 56(5): 542-547). It has also been revealed that many inflammation-associated genes, including IL-2, IL-6 and G-CSF, are varied in their expression levels in response to the signal through the common pathway (Trends Genet. 1999, 15(6): 229-235). A survey was performed by using ATAC-PCR method (adaptor-competitive PCR method: Nucleic Acids Res. 1997, Nov 15; 25(22): 4694-6) for genes of which expression levels were varied depending on stimulation of inflammatory cytokines, TNF- and IL-1. It is possible that genes of which expression is varied in response to this stimulation also participate in inflammation.

[0273] Jurkat cells (Dainippon Pharmaceutical Co., Ltd.: catalog No. 06-152) were cultured in a PRMI1640 medium (Nikken Biological and Medical Institute: catalog No. 14-501F) containing 10% fetal calf serum until the cell count reached  $10^7$  cells. The cells were transferred into a fresh medium containing 10 ng/ml TNF- (recombinant Tumor Necrosis Factor; Wako pure chemical Industries Inc.: catalog No. 201-13461) or IL-1. (recombinant Interleukin-1.; PeprotechEC: catalog No. 200-01B) and, further, cultured at 37. under an atmosphere of 5% CO<sub>2</sub>. The cells cultured in the presence of TNF- were harvested 1, 3 and 7 hours after addition of TNF-. The cells cultured in the presence of IL-1. were harvested 1 and 7 hours after addition of IL-1.. Total RNA was extracted from each of the cells by AGPC method (Acid-Guanidinium-Phenol-Chloroform method: Ana Biochem. 1987, Apr; 162(1):156-9). Total RNA was also extracted from the cells in the absence of any stimulation of TNF- and IL-1

[0274] ATAC-PCR analysis is performed basically according to the same procedure as described in "DNA Microarray and Advanced PCR Methods" (Cell Engineering, p. 104-112, (additional volume, Genome Science Series 1), Muramatsu & Naba (eds.), Shujunnsya). Adaptor ligation reaction was performed for an internal standard sample (which was used for preparing a calibration curve for the assessment of the test samples) and test samples in the following independent two reaction systems. Combinations of each type of the 6 adaptors (AD-1, AD-2, AD-3, AD-4, AD-5, and AD-6: see the sequences shown below) with each sample are as follows:

### Reaction system A

AD1: internal standard sample (x10 concentration)  
AD2: sample before stimulation  
AD3: internal standard sample (x3 concentration)  
AD4: sample with IL-1 stimulation for 1 hour  
AD5: sample with IL-1 stimulation for 7 hours  
AD6: internal standard sample (x1 concentration)

### Reaction system B

AD1: internal standard sample (x1 concentration)  
AD2: sample with TNF stimulation for 1 hour  
AD3: sample with TNF stimulation for 3 hours  
AD4: internal standard sample (x3 concentration)  
AD5: sample with TNF stimulation for 7 hours  
AD6: internal standard sample (x10 concentration)

### Adaptor sequence

AD1;

SEQ ID NO:4180//5'-GTACATATTGTCGTTAGAACCGG-3'

SEQ ID NO:4181//3'-CATGTATAACAGCAATCTTGGCCTAG-5'

AD2;

SEQ ID NO:4182//5'-GTACATATTGTCGTTAGAACGCGACT-3'

SEQ ID NO:4183//3'-CATGTATAACAGCAATCTTGGCTGACTAG-5'

AD3;

SEQ ID NO:4184//5'-GTACATATTGTCGTTAGAACGCGCATACT-3'

SEQ ID NO:4185//3'-CATGTATAACAGCAATCTTGGCGTATCACTAG-5'

AD4;

SEQ ID NO:4186//5'-GTACATATTGTCGTTAGAACGCGATCCATACT-3'

SEQ ID NO:4187//3'-CATGTATAACAGCAATCTTGGCTAGGTATGACTAG-5'

AD5;

SEQ ID NO:4188//5'-GTACATATTGTCGTTAGAACGCGTCAATCCATACT-3'

SEQ ID NO:4189//3'-CATGTATAACAGCAATCTTGGCAGTTAGGTATGACTAG-5'

AD6;

SEQ ID NO:4190//5'-GTACATATTGTCGTTAGAACGCGTACTCAATCCATACT-3'

SEQ ID NO:4191//3'-CATGTATAACAGCAATCTTGGCATGAGTTAGGTATGACTAG-5'

**[0275]** In this assay, the internal standard samples used were total RNA from cultured cells or human tissues from which the cDNA libraries originated. The cultured cells and the total RNAs from tissues are indicated below. Culture of the cells was performed according to the method as described in the supplier's instruction manual. RNA preparation was carried out by standard methods.

Human teratocarcinoma cell NT-2 (Stratagene, catalog No. 204101)

Human neuroblastoma cell SK-N-MC (Dainippon Pharmaceutical Co., Ltd., catalog No. 04-010)

Human neuroblastoma cell Y79 (Dainippon Pharmaceutical Co., Ltd., catalog No. 04-018)

Human placenta tissues total RNA (BioChin, catalog No. 064008)

Human breast tissue total RNA (Clontech, catalog No. 64037-1)

**[0276]** PCR primers used for amplification of specific genes, and names of the corresponding cDNA clones are shown below. The assay was not carried out for clones of which corresponding internal standard sample could not be prepared for the assay. The gene-specific primers were designed so that the PCR products derived from the cDNAs with adaptor were 70-200 bp in size. Sequence of the adaptor-specific primer (labeled with fluorescent dye (FAM)) used for the competitive PCR was GTACATATTGTCGTTAGAACGC (22 nucleotides, SEQ ID NO: 4192). PCR was performed basically at 94. for 5 minutes; and at 94. for 30 seconds, at 50. for 60 seconds, and at 72. for 60 seconds for 30 cycles. The annealing temperature was, however, changed in some PCR experiments.

**[0277]** Nucleotide sequence of clone-specific primer (all the primers consist of 20 nucleotides) used in this experiment

**[0278]** Clone names, primer sequences, and SEQ ID NOs were shown in this order, separated with a double-slash mark, //

5 Y79AA1000967//GGCACAGACACCATCCTTGA//SEQ ID NO:4461  
 Y79AA1001056//ACAAATGAGCCTGAAAAGTC//SEQ ID NO:4462  
 Y79AA1001062//TGGTCTCACTGCCTTCAA//SEQ ID NO:4463  
 Y79AA1001090//AGTCCCCTCAAAGCTCCAGT//SEQ ID NO:4464  
 Y79AA1001212//ACGAAAGCACTCAAATGTCA//SEQ ID NO:4465  
 10 Y79AA1001272//GAATGAAATGTGTTGAGCA//SEQ ID NO:4466  
 Y79AA1001426//AATGATTCGGGGCAGCAGGA//SEQ ID NO:4467  
 Y79AA1001427//GAGAGAGACACACACAAA//SEQ ID NO:4468  
 15 Y79AA1001523//AGTTTTATACCAGCATTGGC//SEQ ID NO:4469  
 Y79AA1001530//GGTGTAGAAGTAAATGGGA//SEQ ID NO:4470  
 Y79AA1001592//GATTGTGTCTCTTACTCCT//SEQ ID NO:4471  
 20 Y79AA1001727//GCTCCACCTGACGTTCTTTA//SEQ ID NO:4472  
 Y79AA1001795//GTCTCCCATATCGCTGTCTT//SEQ ID NO:4473  
 Y79AA1001803//CACTTTCTAATAACCCCTGG//SEQ ID NO:4474  
 25 Y79AA1001863//TTGGGATTGGAACCCGATT//SEQ ID NO:4475  
 Y79AA1001874//AGAAACCACTGAGGCCCAAG//SEQ ID NO:4476  
 Y79AA1002058//CAGAAGCAGAAGCAGGAGCA//SEQ ID NO:4477  
 Y79AA1002121//ATTACTCGGATTCTCCTG//SEQ ID NO:4478  
 30 Y79AA1002129//GAGTTTCTTTGCTAGTTCCA//SEQ ID NO:4479  
 Y79AA1002334//ATATTTGTGTTGCCCTGGG//SEQ ID NO:4480  
 Y79AA1002373//GGATGGCTGGGTCAAATGGT//SEQ ID NO:4481  
 35 Y79AA1002376//AATGATGGCTAGGGTGACTT//SEQ ID NO:4482  
 Y79AA1002378//TCTTCCACATTCGTTACAC//SEQ ID NO:4483  
 Y79AA1002381//AGGGAGTAGATGTTGGTAAA//SEQ ID NO:4484  
 40

[0279] The result of expression frequency analysis is shown in Table 367. Only clones with correlation coefficient of 0.9 or higher are indicated in this Table. Clones that are not presented in the Table include clones for which the assay could not performed because of low expression levels thereof in internal standard samples or because of unexpectedly smaller or larger sizes of the PCR products.

[0280] Among the clones that could be analyzed, clones of which expression levels increased by two fold in response to the IL-1. stimulation 1 or 7 hours after the stimulation are: NT2RM2000514, NT2RP3001159, MAMMA1001237 and MAMMA1000614.

[0281] Clones of which expression levels increased by two fold in response to the TNF-stimulation 1, 3 or 7 hours after the stimulation are:

NT2RM2000582, NT2RM2002109, NT2RP1000679, NT2RP2003664, NT2RP2005597, NT2RP2004108, NT2RP3001592, NT2RP3002738, NT2RP3004133, NT2RP3004321, NT2RP3004557, NT2RP3004294, MAMMA1001237, MAMMA1000141, MAMMA1000788, MAMMA1002070, PLACE1002547, PLACE1003573, PLACE1004305, PLACE1008744, PLACE1011181, PLACE1010713, PLACE1010011, Y79AA1000776, Y79AA1002129,

[0282] Among the clones of which expression levels increased in response to IL-1. stimulation, MAMMA1001237 was a clone of which expression level was varied in response to TNF- stimulation. Among clones showing higher expression levels (with relative value of 5 or higher) prior to the stimulation, PLACE1002080 is an example of clones



of which expression was suppressed by the stimulation. The expression of the clone decreased by three or more fold in response to the stimulation. These genes were found to be associated with inflammatory reaction induced by IL-1 or TNF-..

5 [0283] In Example 15, the genes of which expression levels were varied by culturing in the presence of TNF-. were analyzed by hybridization with high-density DNA filter. As for 3 clones (NT2RP3004557, NT2RP3004294 and PLACE 1002547), the results obtained by ATAC-PCR method were similar to those obtained by hybridization method. However, the results obtained by ATAC-PCR method were not necessarily consistent with those obtained by the hybridization method. Possible reasons for the inconsistency are the difference in cells used between the two experiments, unavail-  
10 ability of some data in the ATAC-PCR experiment, and the difference in the method of data treatment.

Table 28

15 Expression of each cDNA in human tissues (The Table also contains clones with no description in Examples)

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Clone name	Heart	Lung	P.gland	Thymus	Brain	Kidney	Liver	Spleen
GAPDH(Cr1)	38.210	32.670	23.820	13.580	11.230	21.120	24.910	22.440
$\beta$ actin(Cr2)	279.280	368.870	111.100	117.500	92.880	114.650	82.990	256.790
ADRG1000005	53.882	23.005	32.749	22.858	26.564	24.940	22.644	27.001
ADRG1000007	94.778	85.185	160.457	67.191	101.768	62.489	67.150	73.543
ADRG1000009	11.141	50.520	10.357	7.177	6.013	5.219	14.272	21.225
ADRG1000011	71.656	24.579	29.358	19.473	24.898	30.747	49.220	22.221
ADRG1000027	36.238	25.252	20.855	7.328	11.196	14.298	19.658	11.288
ADRG1000058	66.209	129.497	55.226	49.241	30.219	55.872	67.027	243.436
ADRG1000069	38.630	23.459	28.991	12.540	27.353	33.633	28.774	20.911
ADRG1000077	97.465	63.656	448.427	83.412	71.108	53.740	67.906	89.439
ADRG1000092	89.423	45.692	55.810	26.033	44.148	73.339	96.037	73.091
ADRG1000099	73.675	24.424	36.128	17.024	25.964	41.391	42.837	29.666
ADRG1000136	141.745	63.974	77.017	24.777	33.549	58.986	295.009	84.985
ADRG1000147	394.563	155.829	271.210	92.899	165.627	251.266	253.420	150.294
ADRG1000159	50.073	25.425	39.296	15.194	16.125	20.040	33.720	23.278
ADRG1000160	69.386	31.051	59.416	20.154	39.799	27.027	47.169	20.716
ADRG1000171	57.047	23.011	43.063	23.860	40.581	59.814	117.055	32.630
ADRG1000181	45.892	18.666	34.476	15.434	34.225	32.962	39.693	16.334
BGG111000015	153.242	42.337	92.865	41.003	45.168	88.524	85.990	73.392
BGG111000016	177.367	94.731	119.688	34.159	30.249	98.806	98.783	39.204
BGG111000017	84.712	32.614	38.131	20.878	18.769	32.340	39.666	20.750
BGG111000022	52.468	20.452	67.167	12.167	11.158	18.241	19.197	11.937
BGG111000031	30.008	17.072	40.883	12.585	13.313	15.525	16.757	13.406
BGG111000042	49.926	36.336	51.176	26.964	43.122	43.770	49.107	38.776
BGG111000046	31.618	26.472	34.182	31.854	12.650	25.784	18.430	25.385
BNGH41000020	5031.103	2993.496	1444.841	537.162	6973.542	6029.124	3350.527	3649.144
BNGH41000025	91.717	35.026	73.901	27.713	30.765	36.523	37.596	47.074
BNGH41000026	176.757	77.439	98.345	35.807	56.991	91.310	75.797	70.241
BNGH41000027	65.029	56.353	25.896	22.494	12.763	23.748	17.836	23.859
BNGH41000035	148.779	66.776	119.727	56.576	60.996	96.959	72.461	64.458
BNGH41000037	79.500	29.611	43.438	18.317	20.857	36.272	27.525	24.771
BNGH41000042	224.484	110.084	168.448	104.351	102.259	125.323	86.783	122.959
BNGH41000048	56.144	32.253	54.063	14.729	27.312	22.435	29.566	28.937
BNGH41000056	67.258	18.694	30.075	15.602	10.072	20.735	16.100	7.642
BNGH41000087	98.262	46.173	77.657	35.329	40.900	50.029	50.841	45.285
BNGH41000091	50.895	16.985	28.392	10.147	5.469	22.794	10.725	12.410
BNGH41000157	69.043	34.730	40.597	18.088	27.072	22.074	25.410	24.950
BNGH41000169	44.850	21.770	28.655	11.403	25.991	28.509	25.634	25.843
BNGH41000181	17.163	15.689	13.948	3.996	9.287	13.139	15.553	16.575
BNGH41000198	81.510	36.250	60.860	20.585	26.929	35.751	31.695	28.325
BNGH41000219	30.302	25.156	22.187	13.757	11.208	15.235	27.285	35.709
BNGH41000229	252.790	65.948	93.499	51.108	92.555	101.245	96.716	78.266
BNGH41000237	85.757	46.997	55.170	26.780	33.764	47.456	37.007	39.131
BNGH41000238	17.744	36.938	42.360	14.922	35.749	42.848	39.238	13.241
BNGH41000243	45.446	23.667	44.798	20.875	10.516	23.918	22.443	27.033
BNGH41000270	60.889	18.651	29.618	10.724	15.979	12.351	19.152	22.314
BRAWH1000004	43.673	28.539	7.640	11.388	19.198	14.903	32.353	23.777
BRAWH1000018	59.409	17.941	102.270	17.107	709.078	25.732	24.214	24.767
BRAWH1000021	104.772	29.951	51.142	21.042	1169.154	55.762	66.754	27.969
BRAWH1000027	152.205	47.310	67.089	32.199	64.521	70.731	79.670	40.928
BRAWH1000029	106.376	49.221	55.840	40.856	59.552	56.487	64.886	100.132
BRAWH1000040	29.419	16.761	31.101	16.622	30.633	18.200	17.998	15.196
BRAWH1000050	161.264	71.786	118.976	51.863	61.542	97.720	81.271	69.194
BRAWH1000051	74.067	34.341	44.047	20.726	30.434	42.055	53.856	24.624
BRAWH1000060	68.789	22.598	35.012	16.493	19.127	38.662	34.923	28.094
BRAWH1000075	17.318	15.898	36.437	8.901	18.133	17.219	9.321	11.200
BRAWH1000081	43.025	12.998	28.267	7.655	123.677	17.673	15.924	9.844
BRAWH1000084	174.384	42.178	80.534	47.752	152.188	77.111	110.167	102.296
BRAWH1000095	118.239	59.676	64.528	28.174	116.975	53.814	746.700	35.985
BRAWH1000096	146.112	44.967	85.882	27.491	145.013	52.880	52.427	58.678
BRAWH1000097	95.841	72.506	174.954	65.637	64.200	73.707	63.827	63.762
BRAWH1000100	11.943	19.037	18.950	13.536	92.145	16.582	16.646	10.218
BRAWH1000101	134.838	57.232	106.632	40.741	96.396	71.642	88.432	57.336

Table 58

	HEM881000638	55.058	47.453	95.751	42.262	25.684	15.056	22.121	28.829
	HEM881000642	179.188	88.317	251.754	80.865	42.468	81.296	37.696	52.009
5	HEM881000643	43.411	25.689	113.037	18.985	11.038	14.245	8.276	18.743
	HEM881000649	27.852	45.202	137.371	34.816	24.496	9.967	11.881	22.322
	HEM881000652	84.942	61.856	126.562	78.131	42.090	36.343	22.852	31.597
	HEM881000655	418.308	73.377	56.858	57.166	32.733	57.424	38.897	44.477
	HEM881000665	16.253	13.954	10.766	20.817	6.796	13.110	7.987	4.458
	HEM881000668	28.587	13.435	14.606	13.788	25.844	15.049	12.549	11.202
10	HEM881000671	239.020	122.952	561.221	119.970	96.244	75.058	66.812	88.267
	HEM881000673	11.633	5.779	14.629	14.904	5.916	4.811	2.141	12.812
	HEM881000679	16.899	7.357	23.438	7.697	1.049	30.246	7.774	7.063
	HEM881000684	188.240	157.754	430.254	128.150	66.411	89.722	49.173	67.832
	HEM881000692	4.978	9.265	11.569	5.085	1.158	3.240	3.421	1.785
	HEM881000693	63.119	40.561	59.522	22.326	25.408	13.898	31.488	20.706
15	HEM881000705	15.560	31.798	122.757	36.451	19.928	11.568	2.839	10.179
	HEM881000706	22.553	13.626	23.777	8.621	11.683	41.509	10.019	7.584
	HEM881000709	74.737	77.864	245.726	50.833	51.093	50.427	37.955	51.357
	HEM881000714	23.726	10.733	6.625	12.298	6.349	9.891	2.142	14.350
	HEM881000725	24.239	9.575	11.437	13.761	12.596	17.372	8.105	16.144
	HEM881000726	86.971	84.395	208.396	65.157	43.881	37.441	22.020	39.067
20	HEM881000729	51.556	25.288	140.931	23.005	27.775	18.629	12.838	14.902
	HEM881000738	39.002	38.955	166.616	42.588	21.380	43.330	7.181	21.192
	HEM881000749	115.917	94.942	454.741	136.454	54.340	39.253	32.933	49.141
	HEM881000763	47.835	25.201	36.488	16.952	21.036	31.919	14.990	12.111
	HEM881000770	30.598	45.410	167.003	32.786	26.482	25.698	18.186	24.127
	HEM881000774	27.168	21.690	33.470	20.937	12.916	22.598	8.092	17.606
25	HEM881000777	246.286	57.131	58.743	31.851	40.345	119.113	81.364	53.990
	HEM881000781	41.945	36.620	34.149	24.543	23.561	16.383	14.371	20.775
	HEM881000788	10.756	10.608	5.481	6.429	2.950	5.995	4.522	4.589
	HEM881000789	28.490	9.620	26.151	16.088	11.640	16.477	7.916	7.672
	HEM881000790	74.318	56.925	185.959	63.749	33.523	24.232	24.414	28.423
	HEM881000794	18.080	17.254	38.876	24.305	7.427	10.338	5.445	9.305
30	HEM881000807	50.070	31.869	22.751	19.865	20.934	27.002	18.350	27.280
	HEM881000809	334.541	42.976	42.300	26.454	9.545	31.526	31.677	44.152
	HEM881000810	189.355	50.676	163.325	33.349	38.994	74.400	45.398	19.262
	HEM881000821	40.710	9.304	21.006	6.841	5.422	15.981	10.835	5.685
	HEM881000822	8.726	3.570	3.541	1.411	7.255	5.519	1.285	1.525
	HEM881000826	68.485	40.348	201.149	68.467	43.204	31.769	32.812	55.367
	HEM881000827	50.671	34.326	108.391	32.945	15.076	25.813	18.713	25.457
35	HEM881000831	38.060	20.466	29.131	12.368	19.990	20.562	25.373	6.415
	HEM881000835	59.181	56.345	127.358	58.150	44.350	35.831	25.687	35.108
	HEM881000840	117.639	63.375	340.802	61.186	48.924	38.995	20.712	30.526
	HEM881000848	98.938	53.024	210.423	42.569	28.984	47.603	29.642	29.431
	HEM881000852	1.827	2.160	0.621	2.559	1.621	1.272	1.364	1.086
	HEM881000857	16.897	16.768	19.951	14.921	12.912	17.270	10.179	14.915
40	HEM881000858	25.634	16.531	8.162	8.209	14.482	12.749	92.823	10.102
	HEM881000867	106.946	56.331	264.748	50.278	36.949	41.202	26.795	29.760
	HEM881000870	68.550	62.423	192.351	52.406	39.303	55.641	23.738	27.427
	HEM881000876	21.813	12.044	24.968	11.314	7.689	10.690	11.143	26.241
	HEM881000881	30.089	16.478	28.345	14.926	18.419	17.763	18.901	20.494
	HEM881000883	11.669	10.263	26.185	6.975	2.780	8.223	2.906	3.540
45	HEM881000887	42.638	32.274	66.780	22.979	31.512	42.842	20.622	22.566
	HEM881000888	20.318	8.193	11.483	5.178	4.073	8.708	6.801	4.342
	HEM881000890	40.795	42.287	112.076	25.031	11.171	23.116	15.491	16.447
	HEM881000893	38.227	10.603	88.306	24.535	14.440	12.863	9.734	17.727
	HEM881000900	23.814	8.709	17.013	9.267	10.928	12.199	14.105	11.108
	HEM881000905	63.589	43.501	37.125	41.367	26.379	29.649	38.699	31.891
50	HEM881000908	42.944	54.674	120.821	34.982	28.838	28.194	15.897	26.230
	HEM881000910	72.960	51.795	161.850	41.050	36.594	37.378	13.612	23.263
	HEM881000913	33.820	35.219	96.448	24.688	12.371	26.067	14.715	19.268
	HEM881000915	1910.513	222.511	693.345	124.825	532.993	1548.228	1159.943	223.176
	HEM881000917	99.638	64.212	310.142	53.316	39.091	34.989	22.324	40.667
	HEM881000927	80.569	11.252	19.448	8.653	21.944	24.546	17.769	17.391
55	HEM881000932	33.128	33.556	95.029	29.041	17.945	21.758	22.973	31.034

[0284] Expression of each cDNA in human tissues (The Table also contains clones without description in Examples)

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Table 185

Expression of each cDNA in human pulmonary arterial endothelial cells cultured in a medium containing bovine serum albumin, glycated bovine serum albumin or advanced glycation endproduct of bovine serum albumin (This table also contains clones without description in Examples).

In the table, EC\_G\_B/EC\_BSA and EC\_A\_B/EC\_BSA represent the ratios EC\_glycated\_BSA/EC\_BSA and EC\_AGE\_BSA/EC\_BSA, respectively.

Clone_name	EC_BSA	EC_glycated_BSA	EC_AGE_BSA	EC_G_B /EC_BSA	EC_A_B /EC_BSA
GAPDH(Cr1)	100.81	134.21	115.16	1.33	1.14
βactin(Cr2)	1101.9	1092.57	997.36	0.99	0.91
ADRGL1000005	26.88	38.27	36.13	1	1
ADRGL1000007	117.89	127.25	133.21	1.08	1.13
ADRGL1000009	29.18	25.65	26.05	1	1
ADRGL1000011	88.9	117.33	142.9	1.32	1.61
ADRGL1000027	33.24	40.53	43.02	1.01	1.08
ADRGL1000058	153.41	208.84	180.05	1.36	1.17
ADRGL1000069	16.8	21.77	29.81	1	1
ADRGL1000077	25.74	24.72	32.86	1	1
ADRGL1000092	84.52	84.15	121.76	1	1.44
ADRGL1000099	76.19	91.53	106.01	1.2	1.39
ADRGL1000136	52.34	44.76	63.06	0.86	1.2
ADRGL1000147	46.08	45.18	52.15	0.98	1.13
ADRGL1000159	31.52	40.24	42.72	1.01	1.07
ADRGL1000160	52.34	60.37	62.29	1.15	1.19
ADRGL1000171	21.46	16.78	25.59	1	1
ADRGL1000181	37.44	45.71	43.65	1.14	1.09
BGGI11000015	52.42	71	65.47	1.35	1.25
BGGI11000016	127.44	122.93	147.57	0.96	1.16
BGGI11000017	25.65	25.74	31.33	1	1
BGGI11000022	32.82	35.19	25.56	1	1
BGGI11000031	44.42	43.8	40.25	0.99	0.91
BGGI11000042	120.38	146.44	165.42	1.22	1.37
BGGI11000046	74.72	58.85	84.95	0.79	1.14
BNGH41000020	4286.08	3584.67	4330.96	0.84	1.01
BNGH41000025	216.67	223.74	257.06	1.03	1.19
BNGH41000026	25.76	28.16	35.52	1	1
BNGH41000027	29.23	23.83	17.86	1	1
BNGH41000035	280.32	238.34	305.66	0.85	1.09
BNGH41000037	59.14	54.86	54.58	0.93	0.92
BNGH41000042	356.1	324.08	411.07	0.91	1.15
BNGH41000048	1201.37	869.03	739.91	0.72	0.62
BNGH41000056	33.94	31.4	40.01	1	1
BNGH41000087	77.58	81.76	91.07	1.05	1.17
BNGH41000091	21.05	21.23	26.82	1	1
BNGH41000157	81.11	57.28	77.46	0.71	0.95
BNGH41000169	21.1	17.59	22.53	1	1
BNGH41000181	63.54	56.92	70.08	0.9	1.1
BNGH41000198	32.53	26.38	34.37	1	1

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	HEMBB1000858	50	36.71	33.69	0.8	0.8
	HEMBB1000867	68.5	85.51	71.5	1.25	1.04
5	HEMBB1000870	72.76	81.83	67.8	1.12	0.93
	HEMBB1000876	43.35	49.98	46.2	1.15	1.07
	HEMBB1000881	39.02	48.73	55.93	1.22	1.4
	HEMBB1000883	27.77	27.83	32.89	1	1
	HEMBB1000887	266.89	206.91	273.83	0.78	1.03
10	HEMBB1000888	17.01	15.62	22.89	1	1
	HEMBB1000890	137.74	161.89	115.78	1.18	0.84
	HEMBB1000893	74.94	56.68	80.9	0.76	1.08
	HEMBB1000900	20.11	30.62	30.2	1	1
	HEMBB1000905	102.51	124.91	83.08	1.22	0.81
15	HEMBB1000908	78.02	96.72	70.96	1.24	0.91
	HEMBB1000910	49.48	58.33	49.88	1.18	1.01
	HEMBB1000913	46.72	50.63	52.5	1.08	1.12
	HEMBB1000915	1235.16	1265.8	1596.28	1.02	1.29
	HEMBB1000917	69.87	84.21	77.72	1.21	1.11
20	HEMBB1000927	15.28	21.63	16.34	1	1
	HEMBB1000932	45.3	54.47	57.04	1.2	1.26
	HEMBB1000933	136.08	158.95	162.12	1.17	1.19
	HEMBB1000936	64.3	53.14	77.64	0.83	1.21
25	HEMBB1000939	68.9	76.02	84.43	1.1	1.23
	HEMBB1000941	33.76	28.93	53.13	1	1.33
	HEMBB1000947	35.6	29.13	35.69	1	1
	HEMBB1000954	28.25	29.37	27.54	1	1
	HEMBB1000959	63.24	88.65	77.46	1.4	1.22
30	HEMBB1000973	25.02	27.91	25.38	1	1
	HEMBB1000975	16.17	19.83	18.15	1	1
	HEMBB1000981	20.67	24.33	27.87	1	1
	HEMBB1000985	42.97	44	49.41	1.02	1.15
	HEMBB1000991	20.6	19.71	27.25	1	1
35	HEMBB1000996	100.51	115.97	153.69	1.15	1.53
	HEMBB1001000	33.13	32.47	31.91	1	1
	HEMBB1001004	14.02	13.99	23.74	1	1
	HEMBB1001008	29.03	29.07	27.37	1	1
	HEMBB1001011	28.26	27.11	28.64	1	1
40	HEMBB1001014	35.72	32.4	51.5	1	1.29
	HEMBB1001020	53.84	59.19	61.08	1.1	1.13
	HEMBB1001024	103.64	109.34	125.21	1.05	1.21
	HEMBB1001026	119.18	76.31	81.91	0.64	0.69
	HEMBB1001037	57.7	74.41	69.67	1.29	1.21
45	HEMBB1001042	14.13	15.47	21.5	1	1
	HEMBB1001046	14.85	32.01	16.89	1	1
	HEMBB1001047	39.94	47.76	56.41	1.19	1.41
	HEMBB1001048	65.35	74.67	76.93	1.14	1.18
50	HEMBB1001051	28.42	37.58	37.86	1	1
	HEMBB1001056	34.89	38.36	38.94	1	1
	HEMBB1001058	39.94	48.58	50.58	1.21	1.26
	HEMBB1001060	22.94	36.58	33.65	1	1
	HEMBB1001063	27.68	28.29	27.17	1	1
55	HEMBB1001068	29.13	24.56	27.82	1	1
	HEMBB1001082	54.37	76.72	82.56	1.41	1.52

	Y79AA1002373	43.96	55.06	28.34	1.25	0.91
	Y79AA1002376	3080.78	3824.05	4481.1	1.24	1.45
5	Y79AA1002378	73.33	93.61	68.22	1.28	0.93
	Y79AA1002381	248.36	288.51	304.13	1.16	1.22
	Y79AA1002388	118.82	135.82	129.37	1.14	1.09
	Y79AA1002399	36.12	30.1	32.87	1	1
10	Y79AA1002407	57.84	42.82	52.54	0.74	0.91
	Y79AA1002413	78.77	81.36	87.31	1.03	1.11
	Y79AA1002416	34.3	30.2	51.99	1	1.3
	Y79AA1002429	67.91	69.81	80.19	1.03	1.18
15	Y79AA1002431	24.66	21.16	23.98	1	1
	Y79AA1002433	27.12	18.11	23.63	1	1
	Y79AA1002445	78.66	54.58	73.75	0.69	0.94
	Y79AA1002461	29.04	24.84	32	1	1
20	Y79AA1002466	882.69	904.65	782.53	1.02	0.89
	Y79AA1002471	53.74	51.26	68.91	0.95	1.28
	Y79AA1002472	121.95	127.4	127.11	1.04	1.04
	Y79AA1002474	53.33	40.85	47.18	0.77	0.88
25	Y79AA1002482	103.36	111.11	116.07	1.07	1.12
	Y79AA1002487	30.92	25.8	32.51	1	1
	Y79AA1002490	101.4	90.92	90.54	0.9	0.89
	Y79AA1002493	107.88	125.54	105.75	1.16	0.98
30	ZRV6C1006278	46.63	30.08	32.23	0.86	0.86

Table 186

Expression of each cDNA in undifferentiated NT2 cells, in NT2 cells cultured in the presence of retinoic acid, or in NT2 cells that were cultured in the presence of retinoic acid and then further cultured in the presence of cell-division inhibitor added (This table also contains clones without description in Examples)

In the table, NT2, NT2\_RA, and NT2\_RA\_INHIB represent untreated NT2 cells, retinoic acid-treated NT2 cells, and retinoic acid/inhibitor-treated NT2 cells, respectively. The assay was performed in triplicate (n=3), and each result was shown in the column of exp.1, exp.2, or exp.3. In addition, "t-test N/R" and "t-test N/I" represent results of test for significance of difference between the untreated cells and the retinoic acid-treated cells, and between the untreated cells and the retinoic acid/inhibitor-treated cells, respectively. The results of the test

are shown in the columns of \*:p<0.05 and \*\*:p<0.01.

Clonc	NT2			NT2 RA			NT2 RA INHIB			ttest	+	ttest	+
	exp.1	exp.2	exp.3	exp.1	exp.2	exp.3	exp.1	exp.2	exp.3	N/R	-	N/I	-
5	GAPDH(Cr1)	3.53	1.08	0.98	2.92	2.49	2.8	1.76	2.59	1.52			
	$\beta$ actin(Cr2)	155.4	118	99.68	148.5	110.7	101.3	114.7	105.8	151.1			
	ADRGL1000005	4.01	2.03	1.55	4.05	3.65	3.6	2.27	2.93	4.24			
	ADRGL1000007	11.08	5.73	7.92	15.42	10.6	13.87	8.99	8.17	9.15			
	ADRGL1000009	1.11	0.72	1.04	1.66	1.89	1.03	1.22	1.62	1.58		*	+
10	ADRGL1000011	4.27	2.7	2.85	4.32	4.35	3.38	2.76	3.27	3.06			
	ADRGL1000027	1.83	0.38	0.56	0.97	0.62	0.99	0.92	1.33	1.5			
	ADRGL1000058	3.65	2.58	1.37	2.92	3.36	2.75	2.25	3.51	2.7			
	ADRGL1000069	3.25	1.85	3.28	1.86	2.53	2.85	2.01	2.89	2.7			
	ADRGL1000077	13.48	10.41	6.71	19.62	17.92	22.59	11.6	16.66	19.34	*	+	
15	ADRGL1000092	5.73	2.8	4.51	7.31	5.01	4.83	3.24	6.16	7.22			
	ADRGL1000099	5.64	3.42	2.08	5.59	3.73	4.24	3.98	3.98	4.06			
	ADRGL1000136	9.97	3.52	4.19	5.77	4.73	5.86	6.61	5.16	5.49			
	ADRGL1000147	23.09	13.85	11.7	14.77	14.96	14.89	17.7	13.3	19.47			
	ADRGL1000159	6.11	2.22	3.37	5.24	2.88	4.15	2.76	2.93	3.59			
20	ADRGL1000160	7.16	3.48	4.19	5.94	4.59	3.41	3.95	4.67	4.25			
	ADRGL1000171	4.84	2.99	3.23	3.52	4.19	4.37	2.55	3.88	3.45			
	ADRGL1000181	5.1	3.65	2.6	3.16	4.06	2.97	2.64	3.06	3.44			
	BGGI11000015	13.95	6.83	6.72	9.61	9.19	10.24	9.94	10.66	10.13			
	BGGI11000016	15.49	5.92	7.09	11.88	11.38	8.72	11.82	10.98	10.51			
	BGGI11000017	7.89	2.99	3.25	4.94	4.94	4.93	3.55	4.27	3.52			
25	BGGI11000022	8.77	5.14	5.91	7.12	7.05	4.54	5.71	5.59	5.9			
	BGGI11000031	4.71	2.16	2.74	4.09	3.29	3.96	4.02	3.67	2.33			
	BGGI11000042	6.37	5.24	3.74	5.63	6.22	4.36	4.66	5.2	4.04			
	BGGI11000046	19.01	12.57	9.23	12.39	15.7	12.37	8.8	10.92	9.17			
	BNGH41000020	859	910.1	603	164	319.2	267.4	638.2	771.6	845.4	**	-	
30	BNGH41000025	5.35	2.06	2.09	2.76	2.76	3.77	4.23	2.01	3.06			
	BNGH41000026	16.2	7.69	7.05	9.34	11.37	9.66	10.13	7.16	10.71			
	BNGH41000027	2.31	2.18	2.5	2.9	3.01	2.82	3.68	3.48	4.21	**	+	**
	BNGH41000035	14.57	8.83	9.36	10.92	9.55	14.75	15.02	15.18	12.2			
	BNGH41000037	10.56	7.46	6.2	8.16	9.21	6.42	3.37	5.45	4.98			
35	BNGH41000042	77.1	50.85	58.45	47.64	53.39	62.67	28.12	35.48	23.44		*	-
	BNGH41000048	3.5	2.19	1.91	4.28	2.87	2.4	1.63	3.01	1.78			
	BNGH41000056	2.57	2.01	1	1.91	2.63	2.15	1.41	2.4	1.79			
	BNGH41000087	9.84	5.84	5.53	12.49	10.24	10.25	11.74	9.68	8.53			
	BNGH41000091	3.37	2.59	1.21	3.29	3.01	1.55	2.95	2.57	2.13			
	BNGH41000157	10.63	5.64	6.15	8.53	9.05	7.74	6.38	6.68	5.75			
40	BNGH41000169	3.77	4.34	3.82	4.9	3.48	3.32	3.4	4.16	4.19			
	BNGH41000181	2.47	1.59	1.84	2.93	2.1	1.8	1.7	2.66	1.59			
	BNGH41000198	8.13	4.64	3.79	5.48	4.35	5.59	4.3	4.15	4.35			
	BNGH41000219	9.61	3.92	4.87	4.17	5.29	5.45	5.24	7.12	7.13			
	BNGH41000229	19.61	13.28	8.68	10.86	11.27	9.36	7.9	9.5	10.85			
45	BNGH41000237	10.9	5.47	6.45	6.65	6.97	7.79	6.36	6.25	5.44			
	BNGH41000238	4.58	7	3.45	5.91	4.68	4.34	4.33	5.44	4.22			
	BNGH41000243	13.85	8.69	8.48	10.19	9.71	8.97	8.23	4.87	5.54			
	BNGH41000270	5.83	2.62	2.35	2.3	3.05	3.44	2.59	3.49	1.3			
	BRAWH1000004	4.19	2.83	2.48	5.04	3.15	3.26	1.44	3.45	2.05			
50	BRAWH1000018	4.85	1.95	2.29	7.47	8.8	8.85	8.68	6.61	7.96	**	+	*
	BRAWH1000021	6.52	5.06	5.87	5.09	6.94	6.44	2.89	6.23	4.28			
	BRAWH1000027	11.64	8.86	7.19	8.24	10.39	11.51	5.58	7.13	8.24			
	BRAWH1000029	9.58	5.15	3.52	6.01	6.72	6	5.08	5.12	5.84			
	BRAWH1000040	4.6	1.89	2.14	2.92	2.71	2.7	2.92	2.5	3.01			
	BRAWH1000050	11.48	4.95	5.19	9.74	7.25	8.62	8.25	8.09	8.93			
55	BRAWH1000051	8.18	3.93	3.19	6.15	5.72	6.02	5.01	4.25	4.44			



Table 221

	HEMBB1000831	5.58	1.72	2.71	4.5	3.81	4.21	2.23	2.64	2.11						
	HEMBB1000835	4	1.57	1.01	4.73	4.53	5.6	3.04	2.52	2.85 *	+					
5	HEMBB1000840	6.38	3.54	3.15	8.28	10.6	8.97	6.91	4.2	4.08 *	+					
	HEMBB1000848	4.7	2.4	2.04	8.23	8.85	8.6	7.06	5.5	6.33 **	+	*		+		
	HEMBB1000852	0.54	0.28	0.27	0.52	0.36	0.24	1.16	0.97	0.61			*		+	
	HEMBB1000857	7.91	6.39	3.23	5.68	6.47	7.09	4.42	3.6	4.37						
	HEMBB1000858	5.33	2.35	2.78	9.3	8.37	8.17	3.94	3.82	2.97 **	+					
10	HEMBB1000867	5.01	2.6	3.3	9.23	10.12	8.69	3.49	5.17	4.45 **	+					
	HEMBB1000870	4.43	1.73	2.81	6.64	6.44	7.5	2.8	3.34	3.99 *	+					
	HEMBB1000876	2.52	1.01	1.78	2.03	2.41	3.32	1.17	1.96	2.6						
	HEMBB1000881	4.52	2.25	2.68	3.85	3.48	4.21	3.8	3.6	3.52						
	HEMBB1000883	1.07	0.87	0.48	2.38	2.52	2.42	1.86	2.24	1.15 **	+					
15	HEMBB1000887	16.17	10.38	8.54	18.39	28.8	26.71	14.31	15.73	15.23 *	+					
	HEMBB1000888	1.52	0.47	0.72	0.71	0.87	1.25	1.08	2.54	2.95						
	HEMBB1000890	4.2	1.91	2.82	6.2	6.22	11.04	3.56	3.57	3.05 *	+					
	HEMBB1000893	3.13	1.95	2.57	3.14	8.44	5.73	3.88	3.35	2.73						
	HEMBB1000900	2.72	1.85	1.78	2.31	2.75	4	1.77	1.83	1.88						
	HEMBB1000905	7.13	4.79	4.05	6.15	5.33	7.36	6.49	7.74	6.04						
20	HEMBB1000908	3.42	1.78	2.53	3.45	3.15	4.99	2.18	3.31	2.95						
	HEMBB1000910	3.27	1.5	0.99	3.5	4.25	4.18	2.64	2.6	2.61 *	+					
	HEMBB1000913	1.53	1.02	1.16	2.35	1.71	3.01	2.43	2.82	3.12			**		+	
	HEMBB1000915	125.5	96.58	90.74	52.7	70.12	78.2	138.4	94.57	151.2 *	-					
	HEMBB1000917	5.94	3.71	3	10.02	9.8	10.14	6.41	5.43	5.2 **	+					
25	HEMBB1000927	3.9	2.3	4.04	2.93	2.18	2.45	3.26	2.61	3.09						
	HEMBB1000932	1.41	0.52	1.78	2.08	2.21	2.86	1.55	1.9	0.46						
	HEMBB1000933	63.34	47.44	31.38	44.11	52.4	49.52	46.54	37.21	45.55						
	HEMBB1000936	7.16	3.79	4.04	4.95	3.87	5.38	3.06	2.19	2.36						
	HEMBB1000939	9.8	5.4	5.5	8.13	8.11	6.88	7.11	4.16	5.78						
30	HEMBB1000941	1.26	1.52	1.91	2.33	1.33	3.43	1.03	2.28	3						
	HEMBB1000947	3.84	2.12	3.17	3.27	3.95	6.16	2.65	3.42	5						
	HEMBB1000954	2.09	0.96	1.77	3.22	2.47	2.01	1.52	2.5	2.09						
	HEMBB1000959	1.47	0.69	1.99	4.15	4.21	5.2	2.08	3.64	2.15 **	+					
	HEMBB1000973	0.93	0.22	1.08	1.36	1.53	1.02	0.58	1.34	0.88						
	HEMBB1000975	6.35	2.45	2.52	2.87	4.55	4.7	3.97	3.56	3.46						
35	HEMBB1000981	1.55	0.65	1.17	2.92	1.74	2.12	1.91	1.15	1.6						
	HEMBB1000985	4.16	2.16	3.38	6.79	6.53	7.43	6.9	5.56	5.46 **	+	*		+		
	HEMBB1000991	2.4	0.94	2.24	1.58	2.01	2.39	1.83	3.86	2.04						
	HEMBB1000996	6.16	2.86	5.71	15.05	12.65	14.03	9.39	6.89	7.92 **	+					
	HEMBB1001000	0.81	0.42	1.96	2.31	1.45	2	2.11	2.4	1.74						
40	HEMBB1001004	0.63	0.42	0.74	2.36	1.33	1.9	1.27	2.5	0.58 *	+					
	HEMBB1001008	0.9	0.72	1.22	1.95	1.11	0.92	0.7	1.72	0.82						
	HEMBB1001011	4.86	1.41	1.32	2.52	2.1	3.78	2.71	1.63	2.77						
	HEMBB1001014	5.41	3.41	2.83	4.86	8.33	8.51	5.54	2.65	5.28						
	HEMBB1001020	3.52	1.22	3.22	5.91	7.22	5.47	4.21	2.46	3.29 *	+					
	HEMBB1001024	3.88	2.55	2.6	4.94	7.97	7.2	4.48	3.54	3.57 *	+					
45	HEMBB1001026	4.57	3.08	2.54	5.25	5.33	6.61	2.93	3.4	3.78 *	+					
	HEMBB1001037	2.04	0.83	2.17	4.63	4.48	3.78	3.41	3.94	2.4 **	+					
	HEMBB1001042	2.63	0.37	1.26	3.42	3.22	3.69	2.16	3.39	1.69 *	+					
	HEMBB1001046	3.55	2.14	2.26	3.89	3.63	3.68	3.15	4.56	3.14						
	HEMBB1001047	5	1.57	1.46	5.39	4.72	4.88	2.39	1.51	4.62						
50	HEMBB1001048	8.53	3.68	3.67	9.65	6.39	8.39	5.59	5.14	7.15						
	HEMBB1001051	1.18	0.9	0.65	0.91	1.6	1.29	0.9	1.3	2.48						
	HEMBB1001056	4.02	2.51	1.82	4.56	3.43	4.23	3.26	2.37	3.48						
	HEMBB1001058	4.62	1.41	2.29	4.81	4.08	5.54	4.01	2.62	3.49						
	HEMBB1001060	1.13	0.14	0.28	1.95	1.91	2.6	0.75	1.53	1.56 *	+					
55	HEMBB1001063	4.1	1.41	1.69	3.82	4.69	5.11	3.01	2.79	2.86						

Table 366

Expression of each cDNA in synovial cells or in the synovial cells in the presence of TNF  
(This table also contains clones without description in Examples)

In the table, Synoviocyte and Synoviocyte\_TNF represent synovial cells and TNF-treated synovial cells, respectively. The assay was performed in triplicate (n=3), and each result is shown in the column of exp.1, exp.2, or exp.3. In addition, "t-test vs TNF" represents a result of test for significance of difference between the untreated synovial cells and the TNF-treated synovial cells. The increase and decrease in the expression level of a particular gene in response to TNF are represented by + and -, respectively. The results of test for significance of difference are shown in the columns of \*:p<0.05 and \*\*:p<0.01.

Clone	Synoviocyte			Synoviocyte_TNF			t test Inc.	
	exp.1	exp.2	exp.3	exp.1	exp.2	exp.3	TNF	Dec.
GAPDH(Cr1)	0.4	0.8	0.89	0.9	1	1.15		
$\beta$ actin(Cr2)	385.94	262.23	582.98	443.28	422.61	573.47		
ADRGL1000005	2.72	2.97	4.46	7.27	7.45	3.51		
ADRGL1000007	4.36	5.19	9.58	20.78	19.59	18.29	**	+
ADRGL1000009	0.99	1.25	1.64	2.16	4.08	2.02		
ADRGL1000011	1.98	3.56	5.24	22.22	23.49	19.81	**	+
ADRGL1000027	0.79	1.22	1.66	2.82	4.99	1.9		
ADRGL1000058	4.12	7.08	26.9	62.55	67.32	49.15	**	+
ADRGL1000069	1.91	1.68	2.47	14.19	14.54	13.74	**	+
ADRGL1000077	1.98	2	2.54	5.5	2.9	4.16		
ADRGL1000092	2.99	4.79	12.53	21.46	22.09	26.19	**	+
ADRGL1000099	2.77	4.79	12.85	23.61	24.02	25.56	**	+

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	HEMBB1000714	1.41	2.85	9.32	5.31	10.37	8.79	
	HEMBB1000725	1.61	2.22	4.35	3.04	6.22	4.72	
	HEMBB1000726	1.88	2.34	8.76	5.63	7.1	4.83	
5	HEMBB1000729	1.82	3.28	4.3	3.3	5.21	2.79	
	HEMBB1000738	1.94	2.6	5.55	3.99	5.53	6.15	
	HEMBB1000749	4.06	4.15	7.47	7.48	9.56	8.27	
	HEMBB1000763	5.81	5.56	6.21	6.65	9.9	6.61	
10	HEMBB1000770	2.76	2.06	8.8	7.73	9.62	8.83	
	HEMBB1000774	1.62	2.75	3.7	3.07	4.39	2.34	
	HEMBB1000777	5.17	5.49	7.42	6.86	4.9	7.37	
	HEMBB1000781	3.7	4.19	6.89	7.64	5.28	6.83	
	HEMBB1000788	0.87	1.79	2.45	2.65	4.88	1.35	
15	HEMBB1000789	1.91	2.22	3.01	3.1	6.55	1.86	
	HEMBB1000790	1.97	2.15	4.48	4.59	4.21	2.53	
	HEMBB1000794	1.46	1.8	2.85	2.97	3.84	2.06	
	HEMBB1000807	2.55	2.72	5.18	3.57	4	4.26	
20	HEMBB1000809	30.31	26.87	132.99	158.22	156.74	195.14	* +
	HEMBB1000810	1.98	2.67	4.51	3.76	6.08	4.12	
	HEMBB1000821	1.98	1.93	2.98	2.05	4.45	1.79	
	HEMBB1000822	1.08	1.97	2.31	1.65	5.31	1.46	
	HEMBB1000826	1.36	1.99	3.57	3.77	6.11	3.6	
25	HEMBB1000827	2.48	2.89	5.83	2.67	5.05	2.99	
	HEMBB1000831	3.4	2.31	5.67	3.84	7.74	2.95	
	HEMBB1000835	1.76	1.94	6.2	7.59	7.62	7.47	* +
	HEMBB1000840	1.27	2.95	6.89	4.48	7.19	3.01	
	HEMBB1000848	2.08	3.45	5.63	5.39	6.45	5.3	
30	HEMBB1000852	1.26	2.16	2.8	1.07	4.51	1.55	
	HEMBB1000857	7.65	6.49	8.13	7.01	10.69	11.53	
	HEMBB1000858	3.7	3.13	7.3	7.07	9.38	7.31	
	HEMBB1000867	2.21	1.84	4.9	3.02	5.55	4.04	
	HEMBB1000870	1.64	2.37	4.56	2.84	5.31	3.63	
35	HEMBB1000876	1.48	2.86	3.91	4.54	3.22	3.93	
	HEMBB1000881	3.35	5.56	10.5	6.12	5.88	3.85	
	HEMBB1000883	1.02	2.68	2.2	3.03	3.32	2.58	
	HEMBB1000887	16.9	14.54	43.41	67.39	61.26	59.84	* +
	HEMBB1000888	1.03	1.67	2.39	1.63	3.92	1.86	
40	HEMBB1000890	2.93	3.36	10.85	6.01	8.62	7.68	
	HEMBB1000893	3.28	2.54	5.46	4.5	6.14	5.57	
	HEMBB1000900	1.27	1.53	2.98	2.06	2.54	1.58	
	HEMBB1000905	5.09	3.75	6.6	10.05	9.45	8.77	** +
	HEMBB1000908	3.34	2.79	3.01	4.48	4.71	5.7	** +
45	HEMBB1000910	1.74	2.91	2.55	2.09	3.56	2.24	
	HEMBB1000913	1.41	1.51	2.22	2.8	3.41	1.91	
	HEMBB1000915	32.08	25.6	50.05	48	58.92	51.07	
	HEMBB1000917	2.1	2.78	5.72	2.99	4.52	3.44	
50	HEMBB1000927	1.45	1.24	1.82	1.49	3.25	1.88	
	HEMBB1000932	0.66	2.06	2.74	1.81	3.41	1.61	
	HEMBB1000933	7.47	7.12	10.71	12.88	12.78	19.19	
	HEMBB1000936	1.44	1.96	2.87	3.75	6.44	3.55	
	HEMBB1000939	7.86	7.14	9.02	15.98	15.3	18.25	** +
55	HEMBB1000941	1.53	1.86	3.17	3.99	4.46	3.52	* +
	HEMBB1000947	3.53	3.34	4.61	4.67	6.8	5.72	

Table 367

Difference in the expression level of each clone in response to TNF. stimulation or IL-1. stimulation

Before stimulation, IL1 1h, and IL1 7h represent relative levels of expression in the absence of the stimulation, 1 hour after the IL-1. stimulation, and 7 hours after the stimulation, respectively. TNF 1h, TNF 3h, and TNF 7h represent relative levels of expression 1 hour after the TNF. stimulation, 3 hours after the stimulation, and 7 hours after the stimulation, respectively. Correlation coefficients 1 and 2 indicate the correlation coefficients in the calibration curves prepared based on the data for the internal standard in reaction systems A and B, respectively.

Clone	IL1		TNF			Correlation			
	before		1h	3h	7h	1	2		
	stimulation	1h	7h	1h	3h	7h	1	2	
NT2RM1000858	5.6	7.6	3.8	4.7	2.1	1.7	0.98	0.94	
NT2RM1000462	0.9	0.9	0.5	0.7	0.1	0	1	1	
NT2RM1000855	1	1.3	1	1.1	0.4	0.4	1	1	
NT2RM1000789	1	0.9	0.4	1	0.4	0.6	0.96	0.98	
NT2RM2000306	0.7	1.1	0.3	1.1	0.3	0.1	1	0.98	
NT2RM2000514	0.2	0.2	0.6	0.2	0.1	0.2	0.98	0.96	
NT2RM2001126	0.5	0	0.4	0.3	0.3	1.2	0.99	0.99	
NT2RM2001902	1.3	1.6	0.6	1.3	0.8	0.8	1	1	
NT2RM2001738	1.6	1.8	1.5	1.7	0.8	0.9	0.98	1	
NT2RM2000582	0.2	0.1	0	0.7	0.1	0.1	0.99	0.99	
NT2RM2000773	1.1	1.2	1.4	2	1	0.8	0.95	1	
NT2RM2001626	0.4	0.2	0.6	0.7	0.1	0.7	1	1	
NT2RM2001643	1.6	3.1	1.2	2.4	0.7	0.8	1	1	
NT2RM2001792	0.2	0	0	0.3	0.1	0.1	0.98	0.97	
NT2RM2000589	0.2	0.1	0	0.1	0	0	1	0.99	
NT2RM2000588	0.6	0.7	0.1	0.8	0.2	0.2	1	1	
NT2RM2002109	0	0	0	0.2	0.1	0	0.99	0.99	
NT2RM4000284	6.5	9.1	4.8	10.1	3.4	3	1	1	
NT2RM4001735	3.8	4.6	2.1	5	1.6	1.4	1	1	
NT2RM4000100	0.5	0.6	0.2	0.5	0.3	0.3	0.95	0.95	
NT2RM4000417	0.2	0	0	0.2	0.1	0	0.99	0.98	
NT2RM4000761	3.2	3.2	2.2	2.6	0.7	0.7	0.95	1	
NT2RM4001843	1.5	1.8	1.7	2.8	1.2	0.6	0.98	1	
NT2RP1000239	2.1	3.2	1.2	2.1	0.5	0.6	1	0.99	
NT2RP1000465	0.9	0.3	0.3	0.9	0.2	0.1	0.97	0.96	
NT2RP1000679	0.3	0.3	0.4	0.9	0.2	0.3	0.97	1	
NT2RP1001031	1.4	1.4	0.4	1.2	0.1	0.3	1	0.98	
NT2RP2001200	2	1.5	0.8	2.2	0.7	0.7	0.99	1	
NT2RP2001562	2.7	2.4	0.7	3.6	0.4	1.1	1	0.94	
NT2RP2001948	1.1	1.5	0.7	1.3	0.6	0.7	0.97	0.99	
NT2RP2002015	1.3	1.7	0.7	1.8	0.6	0.5	0.99	1	
NT2RP2003390	2	1.7	1.3	2.3	0.6	0.5	0.99	0.99	
NT2RP2003664	0.4	0.1	0.1	0.8	0.1	0	0.99	0.99	
NT2RP2005597	1.2	1.4	0.5	2.7	2.2	2.2	0.96	0.99	
NT2RP2001469	1.7	1.4	1.2	2	0.6	0.6	1	1	
NT2RP2000240	0.9	0.9	0.3	1.4	0.7	0.3	1	1	
NT2RP2000610	2.4	2.2	2.1	2.7	1.5	1.6	0.93	0.96	

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	NT2RP2001276	1	0.4	0.4	0.8	0.2	0.7	0.95	1
	NT2RP2001817	1.2	0.8	0.5	1.9	0.7	0.7	1	1
5	NT2RP2004069	0.6	0.6	0.4	0.8	0.5	0.3	0.93	0.97
	NT2RP2004108	0.3	0.2	0.6	1.1	0.4	0.5	0.96	1
	NT2RP2005391	0.7	0.5	0.1	1.2	0.3	0.3	1	0.99
	NT2RP2006092	1.6	1.2	0.9	2.1	0.6	0.7	0.97	1
	NT2RP2006134	1.2	1.5	0.7	1.9	1	0	0.91	1
10	NT2RP2000818	0.9	0.3	0.3	1.6	0.3	0.3	0.95	1
	NT2RP2000092	1.8	1.8	0.8	2	1	1	0.99	0.98
	NT2RP2000092	1.1	1.1	0.5	1.4	0.6	0.6	0.99	0.97
	NT2RP2001538	2.1	1.9	1.8	2.5	0.6	0.8	0.98	1
	NT2RP2006476	2.1	2.2	1.4	3.2	1.6	2	0.97	0.98
15	NT2RP3000616	0.1	0.1	0	0	0	0	1	1
	NT2RP3000721	2.2	2.8	0.7	2.4	0.4	0.4	1	0.98
	NT2RP3001044	1.5	1.9	0.6	2	0.7	0.4	1	1
	NT2RP3001240	0.8	1	0.8	1.5	0.6	0.7	0.97	0.99
	NT2RP3001592	0.3	0.8	0.8	1.1	0	0	0.94	0.93
20	NT2RP3002448	4.6	4.2	2.5	4.5	0.8	1.2	1	0.98
	NT2RP3002721	1.3	1.6	0.5	1.4	0.3	0.3	1	0.99
	NT2RP3002738	0.1	0	0.1	1.9	0.1	0.1	0.99	1
	NT2RP3002790	1.6	2	0.6	1.7	0.6	0.5	0.98	1
	NT2RP3002836	1.7	3	0.9	2.4	1.6	0.7	1	1
25	NT2RP3003354	0.9	0.7	0.5	0.6	0.4	0.5	0.99	0.92
	NT2RP3003614	0.5	0.4	0	0.3	0.3	0.2	0.99	0.99
	NT2RP3004075	0.8	1.4	0.7	1	0.4	0.4	1	1
	NT2RP3004130	0.3	0.4	0	0.2	0.1	0	0.93	0.96
	NT2RP3004133	1.9	3.5	0.6	3.8	1	1.3	0.99	1
30	NT2RP3004321	0.2	0.2	0	1.4	0.4	0.2	1	0.99
	NT2RP3004406	1.3	0.2	0.2	0.7	0.1	0	1	1
	NT2RP3004552	0.1	0.1	0.1	0.1	0	0	1	1
	NT2RP3004557	1.3	1.1	2.2	2.6	1.5	1.4	0.98	0.94
	NT2RP3004647	1.2	2.1	0.6	1.2	1	0.5	1	1
35	NT2RP3000201	2.3	2.9	0.4	1	1.3	0.5	1	0.98
	NT2RP3000820	1.2	1.6	0.9	1.2	0.6	0.5	1	1
	NT2RP3000818	1.4	1.5	0.7	1.8	0.5	0.7	1	0.99
	NT2RP3001159	1.2	2.5	1.2	1.4	0.6	0.7	0.99	0.99
	NT2RP3002281	1.6	2	1.2	1.8	1	1.2	0.99	1
40	NT2RP3002571	3.9	1.8	1.2	5.2	1.4	0.8	0.99	0.97
	NT2RP3002983	1.4	1.7	0.5	1.4	0.4	0.3	1	1
	NT2RP3003473	0.8	0.9	1	0.7	0.4	0.5	1	0.99
	NT2RP3001976	0.6	1.1	0.1	0.7	0.4	0.1	1	0.99
	NT2RP3002286	1.4	1.8	1	1.6	0.6	0.5	1	0.99
45	NT2RP3002353	7.7	6.4	2.2	8.7	1.1	1.3	0.94	0.99
	NT2RP3004025	1.9	2	1	2.1	1	1	0.96	0.98
	NT2RP3004119	0.8	1.1	0.4	0	0	0.2	1	0.99
	NT2RP3000171	0.7	1.3	0.6	1	0.4	0.3	0.99	1
50	NT2RP3000676	1.2	1.9	0.7	1.1	1.3	0.5	0.99	1
	NT2RP3000921	0.2	0.1	0	0.2	0.1	0	1	0.99
	NT2RP3002015	0.8	0.6	0.4	0.7	0.1	0.1	0.99	0.99
	NT2RP3004294	0	0	0	0.1	0.1	0	1	1
	NT2RP3004345	0.6	0.4	0.2	0.9	0.2	0.5	1	1
55	NT2RP3000148	1.7	2.5	0.8	2	0.8	0.8	1	1

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	NT2RP3000232	0.6	0.8	0.4	0.3	0	0	1	0.99
	NT2RP3001650	2.3	1.5	1.6	1.7	1	1.3	1	1
5	NT2RP3002411	0.5	0.4	0.1	0.5	0.2	0.1	1	1
	NT2RP4001001	0.8	1.3	0.7	1	0.7	0.4	0.97	0.97
	NT2RP4001877	1.5	0.9	1.1	1.2	0.5	0.7	1	0.99
	NT2RP4002451	0.7	1	0.6	0.7	0.2	0.3	0.91	0.95
	NT2RP4000634	1	1	0.3	0.9	0.3	0.4	0.99	1
10	NT2RP4002187	0.4	0.4	0.1	0.7	0.3	0.2	1	0.99
	NT2RP4002715	1.5	1.6	0.7	1.5	0.4	0.3	1	0.99
	MAMMA1000986	3.9	4.1	1.9	4.2	1.8	1.4	0.99	1
	MAMMA1001237	0	0	1.6	0.2	0	0	0.99	0.98
	MAMMA1001978	3.5	3.4	2.3	6	3.4	2.5	0.97	0.98
15	MAMMA1002080	0.4	0	0	0.4	0.1	0	1	0.99
	MAMMA1002234	4	4.4	3	7.7	1.9	3	0.97	1
	MAMMA1000614	4.8	1	15.5	5.6	3.9	4.8	0.95	0.93
	MAMMA1000141	7.1	11.5	3.5	14.8	6.5	3.7	1	0.98
	MAMMA1000706	7.2	9.3	3.9	3.7	2.3	2.6	0.98	0.99
20	MAMMA1000788	3	3.8	2.8	8.9	4.8	4.2	0.92	0.98
	MAMMA1000994	0.3	0	0	0.4	0	0	1	1
	MAMMA1001310	4.1	6.1	3.8	8	2.5	3.6	0.99	0.95
	MAMMA1001344	2.7	4.4	2.2	3.2	2.6	2.1	1	0.99
	MAMMA1001957	2.3	2.7	1.9	1.7	1	1.8	0.99	1
25	MAMMA1002070	0.1	0.1	0	0.8	0.4	0.2	1	0.99
	MAMMA1002586	1.7	1.6	1.2	1.3	0.4	0.3	0.94	1
	MAMMA1000102	2.1	2.3	1.4	3.3	1.6	1.6	1	1
	MAMMA1001066	2.8	2.6	1.8	5.3	0.7	1.2	1	0.98
	MAMMA1001094	2.3	2.9	2	3.3	2.1	2.5	0.96	0.9
30	MAMMA1001609	2	3	1.2	2.7	1.7	2.2	0.99	0.97
	PLACE1002547	2	1.7	1.2	4.1	1.2	2	0.95	1
	PLACE1003573	0	0	0	0.1	0	0	1	0.98
	PLACE1004199	0.1	0.2	0	0	0	0	0.99	0.97
	PLACE1004305	0	0	0	0.3	0	0.2	0.96	0.99
35	PLACE1004450	0.9	0.3	0	0.1	0	0	0.98	0.98
	PLACE1005031	0.9	0	0	0.5	0	0	0.98	0.99
	PLACE1007845	0.8	1	0.4	0.4	0.1	0.1	1	0.98
	PLACE1008984	1.4	1.2	0.4	1.9	0.6	0.5	0.98	0.98
	PLACE1011116	2.6	1.5	1.6	1.6	0.3	0.4	1	1
40	PLACE1000986	0.6	0.2	0.2	0.3	0.1	0.1	1	0.98
	PLACE1004492	1.9	1.9	1.5	3.3	1	1	1	0.97
	PLACE1005569	2.6	0.4	0	1.1	0.3	0.1	0.98	0.99
	PLACE1005601	1.7	1.3	1	2.3	0.6	0.3	0.93	1
	PLACE1006079	0.6	0.3	0	0.1	0.1	0	0.98	0.99
45	PLACE1007077	1.1	0	0	0.3	0.1	0	0.97	0.98
	PLACE1008744	0.4	0.1	0.1	1.1	0.1	0	0.98	1
	PLACE1011181	0.6	0.3	0.5	1.6	0.3	0.5	0.98	0.99
	PLACE1005539	0.4	0	0.2	0.3	0.2	0	1	0.93
	PLACE1008282	1.1	0.7	0.6	1.2	0.4	0.4	0.98	1
50	PLACE1010713	0.6	0.7	0	1.4	0.5	0.4	0.99	0.95
	PLACE1010011	1.2	1.4	0.2	2.7	1.5	1.7	1	0.99
	PLACE3000213	1.9	0.2	0.1	0.8	0.1	0	0.99	1
	PLACE1002080	6.7	3.9	0.3	1.7	0.8	0.5	0.95	0.98
55	SKNMC1000082	1.3	0.1	1.1	0.7	0	0	1	1

	Y79AA1000127	1.8	1.8	1.1	2.1	0.5	0.6	1	1
	Y79AA1000226	1.4	0.8	0.6	0.9	0.3	0.4	0.99	0.99
5	Y79AA1000776	0.3	0.1	0	1.1	0.3	0.5	0.99	0.99
	Y79AA1000876	1.1	1.5	1.2	1.3	0.5	0.8	0.97	1
	Y79AA1001056	1.7	1.7	0.8	1.4	0.9	0.7	1	1
	Y79AA1000777	3.1	3.1	1.2	3.8	0.7	0.5	0.98	0.99
	Y79AA1000030	1	1.3	0.2	1.3	0	0.6	0.98	0.96
10	Y79AA1001212	1.5	1.2	1	2	0.8	0.5	1	0.99
	Y79AA1001427	2.3	3	0.6	2	0.8	0.4	1	1
	Y79AA1001530	0.9	0.9	0.5	1.1	0.4	0.4	1	1
	Y79AA1001592	0.6	0.2	0	0.7	0	0	0.97	1
15	Y79AA1001727	0.8	0.4	0.2	0.9	0.2	0.1	1	1
	Y79AA1001803	0.1	0	0	0.2	0.1	0	0.97	0.99
	Y79AA1002373	0	0	0	0	0	0	0.99	1
	Y79AA1002376	0.9	0.1	0	1.2	0.1	0.4	0.98	1
20	Y79AA1001523	0.5	0.5	0.3	0.6	0.3	0.1	1	0.98
	Y79AA1000888	1.1	1	0.7	1.4	0.7	0.5	1	1
	Y79AA1002129	0.2	0.2	0.1	0.5	0.2	0.2	0.99	1

25 [0285] The present invention has provided a total of 830 novel full length cDNA clones. As has not yet proceeded the isolation of full length cDNA within the human, the invention has a large significance. Those proteins such as secretory proteins, membrane proteins, and proteins associated with signal transduction, glycoprotein, and transcription are known to be associated with many diseases. Those genes and proteins associating with diseases are useful for developing medicines as they can be used as a diagnostic marker, or a target for gene therapy or developing medicines that is capable of regulating their expression and activity. Especially, the cDNA clones encoding a secretion protein are extremely important for medicinal industry since the protein itself is expected to be effective as a medicine, and also the gene may have potential to be associating with many diseases. Moreover, those proteins such as membrane proteins, and proteins associated with signal transduction, glycoprotein, transcription, and diseases, and the genes encoding the proteins may be used as a disease marker. These cDNA clones are also important for medicinal industry as they may be effective for treating diseases through the regulation of the expression and activity of their encoded proteins.

Table 368

40 The names of the representative sequences of the clusters (groups) and the corresponding SEQ IDs.

	HRIFA000016a : 1573	HRIFA017855a : 1979
	HRIFA000071a : 1574	HRIFA017921a : 1980
45	HRIFA000116a : 1575	HRIFA018075a : 1981
	HRIFA000123a : 1576	HRIFA018092a : 1982
	HRIFA000264a : 1577	HRIFA018131a : 1983
	HRIFA000284a : 1578	HRIFA018134a : 1984
50	HRIFA000327a : 1579	HRIFA018238a : 1985
	HRIFA000415a : 1580	HRIFA018262a : 1986
	HRIFA000432a : 1581	HRIFA018287a : 1987
	HRIFA000446a : 1582	HRIFA018447a : 1988
	HRIFA000553a : 1583	HRIFA018580a : 1989
55	HRIFA000564a : 1584	HRIFA018666a : 1990
	HRIFA000631a : 1585	HRIFA018688a : 1991

	HRIFA012333a : 1846	HRIFA028501a : 2252
	HRIFA012354a : 1847	HRIFA028511a : 2253
	HRIFA012417a : 1848	HRIFA028573a : 2254
5	HRIFA012427a : 1849	HRIFA028576a : 2255
	HRIFA012436a : 1850	HRIFA028592a : 2256
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	HRIFA012513a : 1852	HRIFA028651a : 2258
	HRIFA012515a : 1853	HRIFA028652a : 2259
10	HRIFA012584a : 1854	HRIFA028654a : 2260
	HRIFA012625a : 1855	HRIFA028708a : 2261
	HRIFA012692a : 1856	HRIFA028790a : 2262
	HRIFA012702a : 1857	HRIFA028804a : 2263
15	HRIFA012737a : 1858	HRIFA028867a : 2264
	HRIFA012761a : 1859	HRIFA028911a : 2265
	HRIFA012795a : 1860	HRIFA028926a : 2266
	HRIFA012881a : 1861	HRIFA028983a : 2267
	HRIFA012885a : 1862	HRIFA029002a : 2268
20	HRIFA012914a : 1863	HRIFA029050a : 2269
	HRIFA012969a : 1864	HRIFA029107a : 2270
	HRIFA012990a : 1865	HRIFA029208a : 2271
	HRIFA012999a : 1866	HRIFA029209a : 2272
	HRIFA013092a : 1867	HRIFA029256a : 2273
25	HRIFA013103a : 1868	HRIFA029263a : 2274
	HRIFA013135a : 1869	HRIFA029274a : 2275
	HRIFA013235a : 1870	HRIFA029278a : 2276
	HRIFA013254a : 1871	HRIFA029285a : 2277
	HRIFA013265a : 1872	HRIFA029317a : 2278
30	HRIFA013276a : 1873	HRIFA029327a : 2279
	HRIFA013279a : 1874	HRIFA029349a : 2280
	HRIFA013288a : 1875	HRIFA029393a : 2281
	HRIFA013376a : 1876	HRIFA029398a : 2282
35	HRIFA013477a : 1877	HRIFA029425a : 2283
	HRIFA013586a : 1878	HRIFA029434a : 2284
	HRIFA013589a : 1879	HRIFA029440a : 2285
	HRIFA013620a : 1880	HRIFA029460a : 2286
	HRIFA013668a : 1881	HRIFA029467a : 2287
40	HRIFA013726a : 1882	HRIFA029508a : 2288
	HRIFA013744a : 1883	HRIFA029511a : 2289
	HRIFA013899a : 1884	HRIFA029577a : 2290
	HRIFA013911a : 1885	HRIFA029602a : 2291
	HRIFA013919a : 1886	HRIFA029649a : 2292
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	HRIFA013980a : 1888	HRIFA029730a : 2294
	HRIFA014006a : 1889	HRIFA029779a : 2295
	HRIFA014024a : 1890	HRIFA029792a : 2296
	HRIFA014056a : 1891	HRIFA029802a : 2297
50	HRIFA014111a : 1892	HRIFA029866a : 2298
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	HRIFA014178a : 1894	HRIFA030025a : 2300
	HRIFA014185a : 1895	HRIFA030045a : 2301
	HRIFA014303a : 1896	HRIFA030097a : 2302
55	HRIFA014336a : 1897	HRIFA030103a : 2303



	HRIFA014396a : 1898	HRIFA030106a : 2304
	HRIFA014397a : 1899	HRIFA030147a : 2305
	HRIFA014417a : 1900	HRIFA030203a : 2306
5	HRIFA014465a : 1901	HRIFA030237a : 2307
	HRIFA014467a : 1902	HRIFA030248a : 2308
	HRIFA014482a : 1903	HRIFA030250a : 2309
	HRIFA014500a : 1904	HRIFA030264a : 2310
	HRIFA014561a : 1905	HRIFA030342a : 2311
10	HRIFA014568a : 1906	HRIFA030370a : 2312
	HRIFA014590a : 1907	HRIFA030371a : 2313
	HRIFA014598a : 1908	HRIFA030381a : 2314
	HRIFA014620a : 1909	HRIFA030385a : 2315
15	HRIFA014621a : 1910	HRIFA030394a : 2316
	HRIFA014688a : 1911	HRIFA030408a : 2317
	HRIFA014692a : 1912	HRIFA030411a : 2318
	HRIFA014702a : 1913	HRIFA030448a : 2319
	HRIFA014819a : 1914	HRIFA030456a : 2320
20	HRIFA014868a : 1915	HRIFA030461a : 2321
	HRIFA014951a : 1916	HRIFA030472a : 2322
	HRIFA014953a : 1917	HRIFA030509a : 2323
	HRIFA014967a : 1918	HRIFA030511a : 2324
	HRIFA015063a : 1919	HRIFA030545a : 2325
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	HRIFA015122a : 1921	HRIFA030599a : 2327
	HRIFA015129a : 1922	HRIFA030629a : 2328
	HRIFA015219a : 1923	HRIFA030642a : 2329
	HRIFA015236a : 1924	HRIFA030662a : 2330
30	HRIFA015246a : 1925	HRIFA030839a : 2331
	HRIFA015351a : 1926	HRIFA030981a : 2332
	HRIFA015409a : 1927	HRIFA031062a : 2333
	HRIFA015423a : 1928	HRIFA031075a : 2334
	HRIFA015443a : 1929	HRIFA031091a : 2335
35	HRIFA015453a : 1930	HRIFA031126a : 2336
	HRIFA015471a : 1931	HRIFA031249a : 2337
	HRIFA015486a : 1932	HRIFA031336a : 2338
	HRIFA015506a : 1933	HRIFA031350a : 2339
40	HRIFA015536a : 1934	HRIFA031395a : 2340
	HRIFA015547a : 1935	HRIFA031397a : 2341
	HRIFA015568a : 1936	HRIFA031438a : 2342
	HRIFA015671a : 1937	HRIFA031472a : 2343
	HRIFA015682a : 1938	HRIFA031510a : 2344
45	HRIFA015756a : 1939	HRIFA031672a : 2345
	HRIFA015764a : 1940	HRIFA031869a : 2346
	HRIFA015802a : 1941	HRIFA031871a : 2347
	HRIFA015811a : 1942	HRIFA031895a : 2348
	HRIFA015902a : 1943	HRIFA031935a : 2349
50	HRIFA015947a : 1944	HRIFA031986a : 2350
	HRIFA015995a : 1945	HRIFA032009a : 2351
	HRIFA016070a : 1946	HRIFA032011a : 2352
	HRIFA016129a : 1947	HRIFA032066a : 2353
	HRIFA016214a : 1948	HRIFA032067a : 2354
55	HRIFA016240a : 1949	HRIFA032070a : 2355

	HRIFA016255a : 1950	HRIFA032073a : 2356
	HRIFA016290a : 1951	HRIFA032079a : 2357
5	HRIFA016430a : 1952	HRIFA032097a : 2358
	HRIFA016599a : 1953	HRIFA032161a : 2359
	HRIFA016623a : 1954	HRIFA032186a : 2360
	HRIFA016639a : 1955	HRIFA032224a : 2361
10	HRIFA016654a : 1956	HRIFA032257a : 2362
	HRIFA016669a : 1957	HRIFA032271a : 2363
	HRIFA016758a : 1958	HRIFA032274a : 2364
	HRIFA016838a : 1959	HRIFA032275a : 2365
15	HRIFA016963a : 1960	HRIFA032360a : 2366
	HRIFA017031a : 1961	HRIFA032389a : 2367
	HRIFA017146a : 1962	HRIFA032433a : 2368
	HRIFA017190a : 1963	HRIFA032453a : 2369
20	HRIFA017257a : 1964	HRIFA032478a : 2370
	HRIFA017295a : 1965	HRIFA032506a : 2371
	HRIFA017312a : 1966	HRIFA032511a : 2372
	HRIFA017456a : 1967	HRIFA032530a : 2373
	HRIFA017457a : 1968	HRIFA032587a : 2374
25	HRIFA017509a : 1969	HRIFA032605a : 2375
	HRIFA017594a : 1970	HRIFA032642a : 2376
	HRIFA017643a : 1971	HRIFA032696a : 2377
	HRIFA017670a : 1972	HRIFA032730a : 2378
30	HRIFA017703a : 1973	HRIFA032820a : 2379
	HRIFA017729a : 1974	HRIFA032984a : 2380
	HRIFA017791a : 1975	HRIFA033349a : 2381
	HRIFA017801a : 1976	HRIFA033718a : 2382
35	HRIFA017818a : 1977	HRIFA034010a : 2383
	HRIFA017836a : 1978	

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Table 369

The names of the internal sequences that are used in the selection of the clones from the representative sequences, and the corresponding SEQ IDs.

---

AA533598	: 2384	HRIFA036799a	: 2463
AI051329	: 2385	HRIFA037138a	: 2464
HRIFA000595a	: 2386	HRIFA037776a	: 2465
HRIFA000667a	: 2387	HRIFA037838a	: 2466
HRIFA000878a	: 2388	HRIRA000001a	: 2467
HRIFA001269a	: 2389	HRIRA000041a	: 2468
HRIFA001283a	: 2390	HRIRA000058a	: 2469
HRIFA002000a	: 2391	HRIRA000260a	: 2470
HRIFA002196a	: 2392	HRIRA000490a	: 2471
HRIFA003583a	: 2393	HRIRA000522a	: 2472
HRIFA005077a	: 2394	HRIRA000553a	: 2473
HRIFA005781a	: 2395	HRIRA000563a	: 2474
HRIFA006216a	: 2396	HRIRA000640a	: 2475
HRIFA006468a	: 2397	HRIRA000725a	: 2476
HRIFA006822a	: 2398	HRIRA000998a	: 2477
HRIFA007048a	: 2399	HRIRA001053a	: 2478
HRIFA007661a	: 2400	HRIRA001314a	: 2479

	HRIFA007777a : 2401	HRIRA001443a : 2480
	HRIFA007997a : 2402	HRIRA001473a : 2481
	HRIFA008312a : 2403	HRIRA001648a : 2482
5	HRIFA009250a : 2404	HRIRA001690a : 2483
	HRIFA009495a : 2405	HRIRA001726a : 2484
	HRIFA009607a : 2406	HRIRA001884a : 2485
	HRIFA009923a : 2407	HRIRA002098a : 2486
10	HRIFA009978a : 2408	HRIRA002100a : 2487
	HRIFA010730a : 2409	HRIRA002155a : 2488
	HRIFA011029a : 2410	HRIRA002307a : 2489
	HRIFA011416a : 2411	HRIRA002442a : 2490
	HRIFA011461a : 2412	HRIRA002446a : 2491
15	HRIFA012670a : 2413	HRIRA002479a : 2492
	HRIFA012717a : 2414	HRIRA002945a : 2493
	HRIFA012802a : 2415	HRIRA003028a : 2494
	HRIFA013357a : 2416	HRIRA003108a : 2495
	HRIFA013484a : 2417	HRIRA003139a : 2496
20	HRIFA015333a : 2418	HRIRA003819a : 2497
	HRIFA015375a : 2419	HRIRA004049a : 2498
	HRIFA015663a : 2420	HRIRA004286a : 2499
	HRIFA016287a : 2421	HRIRA004583a : 2500
25	HRIFA016302a : 2422	HRIRA004691a : 2501
	HRIFA016782a : 2423	HRIRA004783a : 2502
	HRIFA018555a : 2424	HRIRA005152a : 2503
	HRIFA019338a : 2425	HRIRA005221a : 2504
	HRIFA020315a : 2426	HRIRA005227a : 2505
30	HRIFA020806a : 2427	HRIRA005305a : 2506
	HRIFA022264a : 2428	HRIRA005563a : 2507
	HRIFA022923a : 2429	HRIRA006263a : 2508
	HRIFA023027a : 2430	HRIRA006324a : 2509
	HRIFA023218a : 2431	HRIRA006517a : 2510
35	HRIFA023363a : 2432	HRIRA006580a : 2511
	HRIFA023434a : 2433	HRIRA007665a : 2512
	HRIFA023444a : 2434	HRIRA007680a : 2513
	HRIFA023551a : 2435	HRIRA008129a : 2514
	HRIFA023558a : 2436	HRIRA008152a : 2515
40	HRIFA023641a : 2437	HRIRA008276a : 2516
	HRIFA023798a : 2438	HRIRA008329a : 2517
	HRIFA024330a : 2439	HRIRA008854a : 2518
	HRIFA024338a : 2440	HRIRA008896a : 2519
	HRIFA024384a : 2441	HRIRA008958a : 2520
45	HRIFA024644a : 2442	HRIRA009551a : 2521
	HRIFA025170a : 2443	HRIRA009828a : 2522
	HRIFA025496a : 2444	HRIRA010472a : 2523
	HRIFA025565a : 2445	HRIRA012442a : 2524
50	HRIFA025651a : 2446	HRIRA012921a : 2525
	HRIFA026224a : 2447	HRIRA013325a : 2526
	HRIFA026729a : 2448	HRIRA013644a : 2527
	HRIFA026925a : 2449	HRIRA013675a : 2528
	HRIFA028501a : 2450	HRIRA013702a : 2529
55	HRIFA029454a : 2451	HRIRA013757a : 2530
	HRIFA030181a : 2452	HRIRA013951a : 2531

	HRIFA032701a : 2453	HRIRA014256a : 2532
	HRIFA032801a : 2454	HRIRA014380a : 2533
5	HRIFA033384a : 2455	HRIRA015831a : 2534
	HRIFA033682a : 2456	HRIRA015904a : 2535
	HRIFA033930a : 2457	HRIRA016124a : 2536
	HRIFA034817a : 2458	HRIRA017071a : 2537
10	HRIFA035409a : 2459	HRIRA018191a : 2538
	HRIFA035542a : 2460	HRIRA020304a : 2539
	HRIFA035577a : 2461	HRIRA000579a : 2540
	<u>HRIFA036630a : 2462</u>	

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[0286] The internal sequences include EST, HRIFA(the representative sequence of the 5'-end), and HRIRA (the representative sequence of the 3'-end).

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Table 370

clone	the name of full-length nucleotide sequences	SEQ ID NO of the full-length nucleotide sequences	SEQ ID NO of the deduced amino acid sequences	
	HEMBA1000006	C-HEMBA1000006	2547	2548
	nnnnnnnnnnnn	C-nnnnnnnnnnnn	nnnn	nnnn
15	HEMBA1000121	C-HEMBA1000121	2551	2552
	HEMBA1000128	C-HEMBA1000128	2553	2554
	HEMBA1000275	C-HEMBA1000275	2555	2556
	HEMBA1000300	C-HEMBA1000300	2557	
20	HEMBA1000349	C-HEMBA1000349	2558	2559
	HEMBA1000443	C-HEMBA1000443	2560	2561
	HEMBA1000590	C-HEMBA1000590	2562	2563
	HEMBA1000634	C-HEMBA1000634	2564	2565
	HEMBA1000713	C-HEMBA1000713	2566	2567
25	HEMBA1000745	C-HEMBA1000745	2568	2569
	HEMBA1000907	C-HEMBA1000907	2570	2571
	HEMBA1000940	C-HEMBA1000940	2572	2573
	HEMBA1000962	C-HEMBA1000962	2574	2575
30	HEMBA1001221	C-HEMBA1001221	2576	2577
	HEMBA1001228	C-HEMBA1001228	2578	2579
	HEMBA1001297	C-HEMBA1001297	2580	
	HEMBA1001390	C-HEMBA1001390	2581	2582
35	HEMBA1001563	C-HEMBA1001563	2583	
	HEMBA1001621	C-HEMBA1001621	2584	2585
	nnnnnnnnnnnn	C-nnnnnnnnnnnn	nnnn	nnnn
	HEMBA1001878	C-HEMBA1001878	2588	2589
40	HEMBA1002131	C-HEMBA1002131	2590	2591
	HEMBA1002163	C-HEMBA1002163	2592	2593
	HEMBA1002164	C-HEMBA1002164	2594	2595
	HEMBA1002167	C-HEMBA1002167	2596	2597
45	HEMBA1002178	C-HEMBA1002178	2598	2599
	nnnnnnnnnnnn	C-nnnnnnnnnnnn	nnnn	nnnn

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	HEMBA1002195	C-HEMBA1002195	2602	2603
	HEMBA1002227	C-HEMBA1002227	2604	2605
	HEMBA1002239	C-HEMBA1002239	2606	
5	HEMBA1002316	C-HEMBA1002316	2607	2608
	HEMBA1002420	C-HEMBA1002420	2609	2610
	HEMBA1002421	C-HEMBA1002421	2611	2612
	HEMBA1002524	C-HEMBA1002524	2613	2614
	HEMBA1002551	C-HEMBA1002551	2615	2616
10	HEMBA1002767	C-HEMBA1002767	2617	2618
	HEMBA1002992	C-HEMBA1002992	2619	2620
	HEMBA1003047	C-HEMBA1003047	2621	2622
	HEMBA1003072	C-HEMBA1003072	2623	2624
	HEMBA1003101	C-HEMBA1003101	2625	2626
15	HEMBA1003230	C-HEMBA1003230	2627	2628
	HEMBA1003294	C-HEMBA1003294	2629	
	HEMBA1003315	C-HEMBA1003315	2630	2631
	HEMBA1003392	C-HEMBA1003392	2632	2633
20	HEMBA1003399	C-HEMBA1003399	2634	2635
	HEMBA1003487	C-HEMBA1003487	2636	2637
	HEMBA1003530	C-HEMBA1003530	2638	2639
	HEMBA1003602	C-HEMBA1003602	2640	2641
	HEMBA1003732	C-HEMBA1003732	2642	2643
25	HEMBA1003945	C-HEMBA1003945	2644	2645
	HEMBA1004110	C-HEMBA1004110	2646	2647
	HEMBA1004250	C-HEMBA1004250	2648	2649
	HEMBA1004391	C-HEMBA1004391	2650	2651
	HEMBA1004444	C-HEMBA1004444	2652	2653
30	HEMBA1004454	C-HEMBA1004454	2654	2655
	HEMBA1004505	C-HEMBA1004505	2656	2657
	HEMBA1004797	C-HEMBA1004797	2658	2659
	HEMBA1004982	C-HEMBA1004982	2660	2661
	HEMBA1005070	C-HEMBA1005070	2662	2663
35	HEMBA1005084	C-HEMBA1005084	2664	2665
	HEMBA1005145	C-HEMBA1005145	2666	2667
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	HEMBA1005449	C-HEMBA1005449	2670	2671
40	HEMBA1005489	C-HEMBA1005489	2672	2673
	HEMBA1005522	C-HEMBA1005522	2674	2675
	HEMBA1005545	C-HEMBA1005545	2676	2677
	HEMBA1005698	C-HEMBA1005698	2678	2679
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45	HEMBA1005929	C-HEMBA1005929	2681	2682
	HEMBA1005945	C-HEMBA1005945	2683	2684
	HEMBA1006016	C-HEMBA1006016	2685	
	HEMBA1006171	C-HEMBA1006171	2686	2687
	HEMBA1006299	C-HEMBA1006299	2688	2689
50	HEMBA1006311	C-HEMBA1006311	2690	2691
	HEMBA1006335	C-HEMBA1006335	2692	2693
	HEMBA1006430	C-HEMBA1006430	2694	2695
	HEMBA1006482	C-HEMBA1006482	2696	2697
	HEMBA1006572	C-HEMBA1006572	2698	2699
55	HEMBA1006707	C-HEMBA1006707	2700	2701

	HEMBA1006724	C-HEMBA1006724	2702	2703
	HEMBA1006902	C-HEMBA1006902	2704	2705
	HEMBA1006916	C-HEMBA1006916	2706	2707
5	HEMBA1006960	C-HEMBA1006960	2708	2709
	HEMBA1007013	C-HEMBA1007013	2710	2711
	HEMBA1007057	C-HEMBA1007057	2712	2713
	HEMBA1007241	C-HEMBA1007241	2714	
	HEMBA1007291	C-HEMBA1007291	2715	2716
10	HEMBA1007332	C-HEMBA1007332	2717	
	HEMBB1000276	C-HEMBB1000276	2718	
	HEMBB1000447	C-HEMBB1000447	2719	2720
	HEMBB1000642	C-HEMBB1000642	2721	
15	HEMBB1000668	C-HEMBB1000668	2722	2723
	HEMBB1000679	C-HEMBB1000679	2724	2725
	HEMBB1000881	C-HEMBB1000881	2726	2727
	HEMBB1000905	C-HEMBB1000905	2728	2729
	HEMBB1001026	C-HEMBB1001026	2730	2731
20	HEMBB1001048	C-HEMBB1001048	2732	2733
	HEMBB1001200	C-HEMBB1001200	2734	
	HEMBB1001407	C-HEMBB1001407	2735	2736
	HEMBB1001530	C-HEMBB1001530	2737	2738
	HEMBB1001573	C-HEMBB1001573	2739	2740
25	nnnnnnnnnnnn C-nnnnnnnnnnnnn	nnnn	nnnn	
	HEMBB1001847	C-HEMBB1001847	2743	2744
	HEMBB1001978	C-HEMBB1001978	2745	2746
	HEMBB1002162	C-HEMBB1002162	2747	2748
	HEMBB1002228	C-HEMBB1002228	2749	
30	HEMBB1002245	C-HEMBB1002245	2750	2751
	HEMBB1002427	C-HEMBB1002427	2752	2753
	HEMBB1002465	C-HEMBB1002465	2754	2755
	HEMBB1002663	C-HEMBB1002663	2756	2757
35	HEMBB1002693	C-HEMBB1002693	2758	2759
	MAMMA1000046	C-MAMMA1000046	2760	
	MAMMA1000118	C-MAMMA1000118	2761	2762
	nnnnnnnnnnnn C-nnnnnnnnnnnnn	nnnn	nnnn	
	MAMMA1000449	C-MAMMA1000449	2765	
40	MAMMA1000457	C-MAMMA1000457	2766	2767
	MAMMA1000591	C-MAMMA1000591	2768	2769
	MAMMA1000681	C-MAMMA1000681	2770	2771
	MAMMA1001043	C-MAMMA1001043	2772	2773
	MAMMA1001893	C-MAMMA1001893	2774	2775
45	NT2RM2000241	C-NT2RM2000241	2776	2777
	NT2RM2000306	C-NT2RM2000306	2778	2779
	NT2RM2000410	C-NT2RM2000410	2780	2781
	NT2RM2000423	C-NT2RM2000423	2782	2783
	NT2RM2000497	C-NT2RM2000497	2784	2785
50	NT2RM2000514	C-NT2RM2000514	2786	2787
	NT2RM2000622	C-NT2RM2000622	2788	2789
	NT2RM2001126	C-NT2RM2001126	2790	2791
	NT2RM2001902	C-NT2RM2001902	2792	2793
	NT2RM2001939	C-NT2RM2001939	2794	2795
55	NT2RM2001941	C-NT2RM2001941	2796	2797



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	NT2RM4000198	C-NT2RM4000198	2798	2799
	NT2RM4000284	C-NT2RM4000284	2800	2801
	NT2RM4000295	C-NT2RM4000295	2802	2803
5	NT2RM4000326	C-NT2RM4000326	2804	2805
	NT2RM4000444	C-NT2RM4000444	2806	2807
	NT2RM4000587	C-NT2RM4000587	2808	2809
	NT2RM4000648	C-NT2RM4000648	2810	2811
10	NT2RM4000997	C-NT2RM4000997	2812	2813
	NT2RM4001321	C-NT2RM4001321	2814	2815
	NT2RM4001325	C-NT2RM4001325	2816	2817
	NT2RM4001735	C-NT2RM4001735	2818	2819
	NT2RM4002352	C-NT2RM4002352	2820	2821
15	NT2RP1000002	C-NT2RP1000002	2822	2823
	NT2RP1000050	C-NT2RP1000050	2824	2825
	NT2RP1000181	C-NT2RP1000181	2826	2827
	NT2RP1000261	C-NT2RP1000261	2828	2829
20	NT2RP1000300	C-NT2RP1000300	2830	2831
	NT2RP1000325	C-NT2RP1000325	2832	2833
	NT2RP1000448	C-NT2RP1000448	2834	2835
	NT2RP1000551	C-NT2RP1000551	2836	2837
	NT2RP1000579	C-NT2RP1000579	2838	2839
25	NT2RP1000613	C-NT2RP1000613	2840	2841
	NT2RP1000903	C-NT2RP1000903	2842	2843
	NT2RP1000981	C-NT2RP1000981	2844	2845
	NT2RP1001004	C-NT2RP1001004	2846	2847
	NT2RP1001020	C-NT2RP1001020	2848	2849
30	NT2RP1001563	C-NT2RP1001563	2850	2851
	NT2RP2000394	C-NT2RP2000394	2852	2853
	NT2RP2000479	C-NT2RP2000479	2854	2855
	NT2RP2000514	C-NT2RP2000514	2856	2857
35	NT2RP2000533	C-NT2RP2000533	2858	2859
	NT2RP2000649	C-NT2RP2000649	2860	2861
	NT2RP2000663	C-NT2RP2000663	2862	2863
	NT2RP2000694	C-NT2RP2000694	2864	2865
	NT2RP2000903	C-NT2RP2000903	2866	2867
40	NT2RP2001480	C-NT2RP2001480	2868	2869
	NT2RP2001495	C-NT2RP2001495	2870	2871
	NT2RP2001514	C-NT2RP2001514	2872	2873
	NT2RP2001529	C-NT2RP2001529	2874	2875
	NT2RP2001769	C-NT2RP2001769	2876	2877
45	NT2RP2001878	C-NT2RP2001878	2878	2879
	NT2RP2001903	C-NT2RP2001903	2880	2881
	NT2RP2001915	C-NT2RP2001915	2882	2883
	NT2RP2001956	C-NT2RP2001956	2884	2885
	NT2RP2002063	C-NT2RP2002063	2886	2887
50	NT2RP2002188	C-NT2RP2002188	2888	2889
	NT2RP2002232	C-NT2RP2002232	2890	2891
	NT2RP2002304	C-NT2RP2002304	2892	2893
	NT2RP2002409	C-NT2RP2002409	2894	2895
	NT2RP2002510	C-NT2RP2002510	2896	2897
55	NT2RP2002527	C-NT2RP2002527	2898	2899
	NT2RP2002533	C-NT2RP2002533	2900	2901

	NT2RP2000092	C-NT2RP2000092	4111	4112
	NT2RP2001538	C-NT2RP2001538	4113	4114
5	NT2RP2001921	C-NT2RP2001921	4115	4116
	NT2RP2003138	C-NT2RP2003138	4117	4118
	NT2RP2003302	C-NT2RP2003302	4119	4120
	NT2RP2003950	C-NT2RP2003950	4121	4122
	NT2RP2005535	C-NT2RP2005535	4123	4124
10	NT2RP2005774	C-NT2RP2005774	4125	4126
	NT2RP3000148	C-NT2RP3000148	4127	4128
	NT2RP3000232	C-NT2RP3000232	4129	4130
	NT2RP3000427	C-NT2RP3000427	4131	
15	NT2RP3000652	C-NT2RP3000652	4132	4133
	NT2RP3001650	C-NT2RP3001650	4134	4135
	NT2RP3002409	C-NT2RP3002409	4136	
	NT2RP3002411	C-NT2RP3002411	4137	4138
20	NT2RP3003448	C-NT2RP3003448	4139	
	NT2RP4002715	C-NT2RP4002715	4140	4141
	OVARC1000307	C-OVARC1000307	4142	4143
	PLACE1000907	C-PLACE1000907	4144	4145
25	PLACE1007081	C-PLACE1007081	4146	4147
	PLACE1010011	C-PLACE1010011	4148	4149
	PLACE3000213	C-PLACE3000213	4150	4151
	PLACE4000354	C-PLACE4000354	4152	4153
	PLACE4000455	C-PLACE4000455	4154	
30	THYRO1000776	C-THYRO1000776	4155	4156
	THYRO1001593	C-THYRO1001593	4157	4158
	Y79AA1000750	C-Y79AA1000750	4159	4160
	Y79AA1000888	C-Y79AA1000888	4161	4162
35	Y79AA1002129	C-Y79AA1002129	4163	4164
	Y79AA1002334	C-Y79AA1002334	4165	4166
	MAMMA1002224	C-MAMMA1002224	4167	
40	NT2RP1000271	C-NT2RP1000271	4168	4169
	NT2RP3000481	C-NT2RP3000481	4170	4171
	NT2RP3004481	C-NT2RP3004481	4172	4173
	HEMBA1006658	C-HEMBA1006658	4174	4175
	NT2RP2006099	C-NT2RP2006099	4176	4177
45	NT2RP2006580	C-NT2RP2006580	4178	4179

## Homology search result 1

50

[0287] The result of the homology search in the SwissProt using the representative sequences of the 5'-ends.

55

Indicated are from the top,  
the name of the representative sequence of the cluster,  
definition of the top hit data,  
the P-value: the length of the sequence used for comparison (nucleotide):similarity (%),  
the organism of which the top hit data is obtained,  
the Accession No. of the top hit data.

[0288] Homology search results of the representative sequences of the 5'-end cluster to the data in SwissProt database are shown only for the representative sequences of the cluster from which clones were selected based on the homology search results.

5 [0289] The P-value is the score which is determined by taking into account the statistic probability of occurrence between the two sequences, and generally low score reflects high similarity. (Altschul, S.F., Gish, W., Miller, W., Myers, E.W. & Lipman, D.J. (1990) "Basic local alignment search tool." J. Mol. Biol. 215:403-410; Gish, W. & States, D.J. (1993) "Identification of protein coding regions by database similarity search." Nature Genet. 3:266-272).

10 HRIFA000016a  
GLYCINE-RICH CELL WALL STRUCTURAL PROTEIN 1.8 PRECURSOR (GRP 1.8).  
9.2e-05:178:32  
PHASEOLUS VULGARIS (KIDNEY BEAN) (FRENCH BEAN).  
P10496

15 HRIFA000071a  
CIRCUMSPOROZOITE PROTEB PRECURSOR (CS).  
5.8e-05:194:29  
PLASMODIUM SIMIUM.  
Q03110

20 HRIFA000116a  
HYPOTHETICAL 68.7 KD PROTEIN ZK757.1 IN CHROMOSOME III.  
6.2e-06:83:27  
CAENORHABDITIS ELEGANS.  
25 P34679

HRIFA000123a  
PATHOGENESIS-RELATED PROTEIN 1 PRECURSOR (PR-1).  
6.2e-08:89:34  
30 ARABIDOPSIS THALIANA (MOUSE-EAR CRESS).  
P33154

HRIFA000264a  
PROCOLLAGEN ALPHA 1(I) CHAIN PRECURSOR.  
35 1.4e-06:231:34  
GALLUS GALLUS (CHICKEN).  
P02457

40 HRIFA000327a  
ATP-BINDING CASSETTE TRANSPORTER 1.  
2.0e-16:238:31  
MUS MUSCULUS (MOUSE).  
P41233

45 HRIFA000415a  
PROLINE-RICH PROTEIN MP-2 PRECURSOR.  
3.6e-06:120:35  
MUS MUSCULUS (MOUSE).  
P05142

50 HRIFA000432a  
PUTATIVE GENERAL NEGATIVE REGULATOR OF TRANSCRIPTION C16C9.04C.  
2.2e-21:86:52  
SCHIZOSACCHAROMYCES POMBE (FISSION YEAST).  
55 Q09818

HRIFA000446a  
HYPOTHETICAL 64.8 KD PROTEIN IN GDI1-COX15 INTERGENIC REGION.

- SUS SCROFA (PIG).  
P36393
- 5 HRIFA029649a  
TRANS-ACTING TRANSCRIPTIONAL PROTEIN ICP0 (VMW118 PROTEIN).  
0.30:99:34  
HERPES SIMPLEX VIRUS (TYPE 2 / STRAIN HG52).  
P28284
- 10 HRIFA029715a  
GROWTH ARREST AND DNA-DAMAGE-INDUCIBLE PROTEIN GADD153 (DNA-DAMAGE INDUCIBLE TRANSCRIPT 3) (DDIT3) (C/EBP-HOMOLOGOUS PROTEIN) (CHOP).  
0.54:95:30  
HOMO SAPIENS (HUMAN).  
15 P35638
- HRIFA029730a  
HISTIDINE-RICH GLYCOPROTEIN PRECURSOR.  
3.8e-05:131:29  
20 PLASMODIUM LOPHURAE.  
P04929
- HRIFA029792a  
PUTATIVE SERINE/THREONINE-PROTEIN KINASE PKWA (EC 2.7.1.-).  
25 9.0e-09:178:30  
THERMOMONOSPORA CURVATA.  
P49695
- HRIFA029802a  
30 TRAM PROTEIN (TRANSLOCATING CHAIN-ASSOCIATING MEMBRANE PROTEIN).  
7.2e-73:204:69  
CANIS FAMILIARIS (DOG).  
Q01685
- 35 HRIFA029866a  
PROTEIN KINASE BYR2 (EC 2.7.1.-) (PROTEIN KINASE STE8) (MAPK KINASE KINASE) (MAPKKK).  
1.2e-27:144:45  
SCHIZOSACCHAROMYCES POMBE (FISSION YEAST).  
P28829
- 40 HRIFA029932a  
F-SPONDIN PRECURSOR.  
9.1e-24:191:37  
XENOPUS LAEVIS (AFRICAN CLAWED FROG).  
45 P35447
- HRIFA030025a  
ENDOSOMAL P24A PROTEIN PRECURSOR (70 KD ENOMEMBRANE PROTEIN) (PHEROMONE ALPHA-FACTOR TRANSPORTER) (ACIDIC 24 KD LATE ENDOCYTIC INTERMEDIATE COMPONENT).  
50 1.0e-11:138:31  
SACCHAROMYCES CEREVISIAE (BAKER'S YEAST).  
P32802
- 55 HRIFA030045a  
SARCALUMENIN PRECURSOR.  
2.4e-20:151:32  
ORYCTOLAGUS CUNICULUS (RABBIT).  
P13666

- HRIFA032642a  
PROLINE-RICH PROTEIN MP-2 PRECURSOR.  
5.0e-05:127:33  
MUS MUSCULUS (MOUSE).  
5 P05142
- HRIFA032696a  
COLLAGEN ALPHA 1(II) CHAIN (FRAGMENTS).  
1.4e-13:200:38  
10 BOS TAURUS (BOVINE).  
P02459
- HRIFA032730a  
K-GLYPICAN PRECURSOR.  
15 4.8e-67:180:68  
MUS MUSCULUS (MOUSE).  
P51655
- HRIFA032820a  
20 GLUTAMIC ACID-RICH PROTEIN PRECURSOR.  
7.5e-05:192:23  
PLASMODIUM FALCIPARUM (ISOLATE FC27 / PAPUA NEW GUINEA).  
P13816
- 25 Homology search result 2
- [0290]** Homology of representative sequences of the 5'-end cluster to the data in Swiss-Prot database  
**[0291]** Representative sequence of the 5'-end cluster exhibiting relatively high homology (304 cluster: "exhibiting relatively high homology" means that the P value is  $10^{-10}$  or less)
- 30 HRIFA000327a, HRIFA000432a, HRIFA000553a, HRIFA000564a, HRIFA000631a, HRIFA000683a, HRIFA000776a, HRIFA000814a, HRIFA001132a, HRIFA001138a, HRIFA001337a, HRIFA001341a, HRIFA001489a, HRIFA001712a, HRIFA001720a, HRIFA001942a, HRIFA001975a, HRIFA001984a, HRIFA002384a, HRIFA002503a, HRIFA002743a, HRIFA002766a, HRIFA002805a, HRIFA002891a, HRIFA002919a, HRIFA002980a, HRIFA003063a, HRIFA003093a, HRIFA003635a, HRIFA004006a, HRIFA004034a, HRIFA004112a, HRIFA004426a, HRIFA004490a, HRIFA004523a, HRIFA004663a, HRIFA004696a, HRIFA004714a, HRIFA004745a, HRIFA004919a, HRIFA005184a, HRIFA005231a, HRIFA005240a, HRIFA005271a, HRIFA005372a, HRIFA005392a, HRIFA005409a, HRIFA005420a, HRIFA005438a, HRIFA005462a, HRIFA005644a, HRIFA005720a, HRIFA005732a, HRIFA005760a, HRIFA005781a, HRIFA006183a, HRIFA006494a, HRIFA006510a, HRIFA006566a, HRIFA006586a, HRIFA006596a, HRIFA006649a, HRIFA006667a, HRIFA006730a, HRIFA006926a, HRIFA007013a, HRIFA007219a, HRIFA007228a, HRIFA007243a, HRIFA007352a, HRIFA007424a, HRIFA007435a, HRIFA007463a, HRIFA007493a, HRIFA007571a, HRIFA007659a, HRIFA007722a, HRIFA007745a, HRIFA008000a, HRIFA008200a, HRIFA008284a, HRIFA008314a, HRIFA008362a, HRIFA008459a, HRIFA008483a, HRIFA008547a, HRIFA008611a, HRIFA008661a, HRIFA008717a, HRIFA008784a, HRIFA008981a, HRIFA009101a, HRIFA009171a, HRIFA009220a, HRIFA009451a, HRIFA009482a, HRIFA009783a, HRIFA009881a, HRIFA010085a, HRIFA010090a, HRIFA010130a, HRIFA010319a, HRIFA010394a, HRIFA010460a, HRIFA010790a, HRIFA010975a, HRIFA011016a, HRIFA011179a, HRIFA011197a, HRIFA011449a, HRIFA011659a, HRIFA011947a, HRIFA012278a, HRIFA012584a, HRIFA012625a, HRIFA012692a, HRIFA012795a, HRIFA012885a, HRIFA012914a, HRIFA012969a, HRIFA012990a, HRIFA013254a, HRIFA013265a, HRIFA013276a, HRIFA013376a, HRIFA013477a, HRIFA013586a, HRIFA013726a, HRIFA013744a, HRIFA013911a, HRIFA014006a, HRIFA014185a, HRIFA014336a, HRIFA014465a, HRIFA014500a, HRIFA014561a, HRIFA014568a, HRIFA014621a, HRIFA014688a, HRIFA014819a, HRIFA014951a, HRIFA014967a, HRIFA015063a, HRIFA015070a, HRIFA015246a, HRIFA015423a, HRIFA015453a, HRIFA015486a, HRIFA015506a, HRIFA015536a, HRIFA015547a, HRIFA015568a, HRIFA015756a, HRIFA015811a, HRIFA016070a, HRIFA016290a, HRIFA016430a, HRIFA016654a, HRIFA016758a, HRIFA017031a, HRIFA017257a, HRIFA017295a, HRIFA017312a, HRIFA017703a, HRIFA017855a, HRIFA018092a, HRIFA018131a, HRIFA018134a, HRIFA018580a, HRIFA018827a, HRIFA018904a, HRIFA018993a, HRIFA019105a, HRIFA019136a, HRIFA019175a, HRIFA019262a, HRIFA019466a, HRIFA019867a, HRIFA019869a, HRIFA020272a, HRIFA020335a, HRIFA020349a, HRIFA020862a, HRIFA021213a, HRIFA021398a, HRIFA021499a, HRIFA021637a, HRIFA021651a, HRIFA021754a, HRIFA021781a, HRIFA022065a, HRIFA022139a, HRIFA022166a, HRIFA022177a, HRIFA022182a, HRIFA022227a, HRIFA022249a, HRIFA022265a,

HRIFA022328a, HRIFA022423a, HRIFA022528a, HRIFA022546a, HRIFA022564a, HRIFA022616a, HRIFA022671a,  
 HRIFA022691a, HRIFA022707a, HRIFA022729a, HRIFA022737a, HRIFA022776a, HRIFA022875a, HRIFA022895a,  
 HRIFA023007a, HRIFA023227a, HRIFA023257a, HRIFA023304a, HRIFA023464a, HRIFA023767a, HRIFA023923a,  
 HRIFA024132a, HRIFA024255a, HRIFA024392a, HRIFA024423a, HRIFA024504a, HRIFA024718a, HRIFA024767a,  
 5 HRIFA024937a, HRIFA024994a, HRIFA025046a, HRIFA025250a, HRIFA025261a, HRIFA025353a, HRIFA025492a,  
 HRIFA025636a, HRIFA025695a, HRIFA025706a, HRIFA025766a, HRIFA025800a, HRIFA025907a, HRIFA025913a,  
 HRIFA026089a, HRIFA026364a, HRIFA026496a, HRIFA026789a, HRIFA026813a, HRIFA026860a, HRIFA027012a,  
 HRIFA027045a, HRIFA027125a, HRIFA027179a, HRIFA027187a, HRIFA027622a, HRIFA027625a, HRIFA027656a,  
 HRIFA027681a, HRIFA027722a, HRIFA027940a, HRIFA028157a, HRIFA028402a, HRIFA028468a, HRIFA028511a,  
 10 HRIFA028651a, HRIFA028790a, HRIFA029002a, HRIFA029208a, HRIFA029209a, HRIFA029256a, HRIFA029263a,  
 HRIFA029285a, HRIFA029317a, HRIFA029327a, HRIFA029393a, HRIFA029511a, HRIFA029802a, HRIFA029866a,  
 HRIFA029932a, HRIFA030025a, HRIFA030045a, HRIFA030250a, HRIFA030342a, HRIFA030370a, HRIFA030371a,  
 HRIFA030411a, HRIFA030448a, HRIFA030545a, HRIFA030629a, HRIFA030642a, HRIFA030662a, HRIFA031336a,  
 HRIFA031869a, HRIFA031986a, HRIFA032009a, HRIFA032011a, HRIFA032070a, HRIFA032073a, HRIFA032079a,  
 15 HRIFA032224a, HRIFA032274a, HRIFA032275a, HRIFA032433a,  
 HRIFA032453a, HRIFA032605a, HRIFA032696a, HRIFA032730a,

## Homology search result 3

20 **[0292]** Representative sequence of the 5'-end cluster exhibiting relatively low homology (221 cluster: "exhibiting  
 relatively low homology" means that the P value is higher than  $10^{-10}$  and  $10^{-4}$  or less)  
 HRIFA000016a, HRIFA000071a, HRIFA000116a, HRIFA000123a, HRIFA000264a, HRIFA000415a, HRIFA000446a,  
 HRIFA000695a, HRIFA000845a, HRIFA001971a, HRIFA002063a, HRIFA002102a, HRIFA002284a, HRIFA002309a,  
 HRIFA002694a, HRIFA002762a, HRIFA002787a, HRIFA003055a, HRIFA003340a, HRIFA003402a, HRIFA003504a,  
 25 HRIFA003892a, HRIFA003946a, HRIFA004162a, HRIFA004401a, HRIFA004780a, HRIFA005072a, HRIFA005102a,  
 HRIFA005214a, HRIFA005255a, HRIFA005300a, HRIFA005369a, HRIFA005702a, HRIFA005728a, HRIFA005944a,  
 HRIFA006298a, HRIFA006448a, HRIFA006572a, HRIFA006633a, HRIFA006642a, HRIFA007068a, HRIFA007244a,  
 HRIFA007262a, HRIFA007512a, HRIFA007532a, HRIFA007565a, HRIFA007728a, HRIFA007909a, HRIFA008174a,  
 HRIFA008426a, HRIFA008596a, HRIFA008790a, HRIFA008989a, HRIFA009578a, HRIFA009825a, HRIFA009852a,  
 30 HRIFA009983a, HRIFA010005a, HRIFA010078a, HRIFA010152a, HRIFA010301a, HRIFA010361a, HRIFA010425a,  
 HRIFA010466a, HRIFA010799a, HRIFA011580a, HRIFA011820a, HRIFA012167a, HRIFA012354a, HRIFA012427a,  
 HRIFA012436a, HRIFA012515a, HRIFA012702a, HRIFA012737a, HRIFA013135a, HRIFA013235a, HRIFA013279a,  
 HRIFA013589a, HRIFA013620a, HRIFA013919a, HRIFA013932a, HRIFA014056a, HRIFA014111a, HRIFA014133a,  
 HRIFA014396a, HRIFA014397a, HRIFA014598a, HRIFA014702a, HRIFA014868a, HRIFA015219a, HRIFA015995a,  
 35 HRIFA016214a, HRIFA016240a, HRIFA016255a, HRIFA016639a, HRIFA016669a, HRIFA016963a, HRIFA017457a,  
 HRIFA017643a, HRIFA017670a,  
 HRIFA017801a, HRIFA017836a, HRIFA017921a, HRIFA018238a, HRIFA018262a, HRIFA018287a, HRIFA018666a,  
 HRIFA018688a, HRIFA018754a, HRIFA018794a, HRIFA018870a, HRIFA018931a, HRIFA019412a, HRIFA019490a,  
 HRIFA019498a, HRIFA019532a, HRIFA019651a, HRIFA020144a, HRIFA020184a, HRIFA020453a, HRIFA020693a,  
 40 HRIFA020707a, HRIFA020748a, HRIFA021061a, HRIFA021224a, HRIFA021494a, HRIFA021794a, HRIFA021855a,  
 HRIFA021906a, HRIFA022156a, HRIFA022203a, HRIFA022234a, HRIFA022702a, HRIFA022728a, HRIFA022782a,  
 HRIFA022865a, HRIFA022890a, HRIFA022985a, HRIFA023048a, HRIFA023069a, HRIFA023129a, HRIFA023154a,  
 HRIFA023212a, HRIFA023489a, HRIFA023634a, HRIFA023894a, HRIFA024088a, HRIFA024197a, HRIFA024218a,  
 HRIFA024473a, HRIFA024482a, HRIFA024543a, HRIFA025327a, HRIFA025479a, HRIFA025488a, HRIFA025703a,  
 45 HRIFA025771a, HRIFA025778a, HRIFA025904a, HRIFA025966a, HRIFA025978a, HRIFA026121a, HRIFA026242a,  
 HRIFA026316a, HRIFA026382a, HRIFA026465a, HRIFA026519a, HRIFA026564a, HRIFA026576a, HRIFA026618a,  
 HRIFA026659a, HRIFA026764a, HRIFA027327a, HRIFA027329a, HRIFA027355a, HRIFA027644a, HRIFA027673a,  
 HRIFA027714a, HRIFA027860a, HRIFA028061a, HRIFA028187a, HRIFA028262a, HRIFA028371a, HRIFA028440a,  
 HRIFA028501a, HRIFA028576a, HRIFA028614a, HRIFA028911a, HRIFA029050a, HRIFA029278a, HRIFA029349a,  
 50 HRIFA029425a, HRIFA029434a, HRIFA029460a, HRIFA029467a, HRIFA029508a, HRIFA029730a, HRIFA029792a,  
 HRIFA030103a, HRIFA030147a,  
 HRIFA030264a, HRIFA030381a, HRIFA030456a, HRIFA030509a, HRIFA030511a, HRIFA030566a, HRIFA030599a,  
 HRIFA031126a, HRIFA031249a, HRIFA031438a, HRIFA031935a, HRIFA032257a, HRIFA032360a, HRIFA032389a,  
 HRIFA032478a, HRIFA032506a, HRIFA032511a, HRIFA032530a, HRIFA032587a, HRIFA032642a, HRIFA032820a,  
 55

## Homology search result 4

**[0293]** Representative sequence of the 5'-end cluster exhibiting low homology (115 cluster: "exhibiting low homology"

P00989

F-HEMBB1000567  
 HISTIDINE-RICH GLYCOPROTEIN PRECURSOR.  
 5.0e-05:131:29  
 PLASMODIUM LOPHURAE.  
 P04929

10 F-HEMBB1000642  
 BASIC PROLINE-RICH PEPTIDE IB-1.  
 0.0074:66:31  
 HOMO SAPIENS (HUMAN).  
 P04281

15 F-HEMBB1000668  
 VEGETATIBLE INCOMPATIBILITY PROTEIN HET-E-1.  
 7.3e-10:184:32  
 PODOSPORA ANSERINA.  
 Q00808

20 F-HEMBB1000679  
 TRAM PROTEIN (TRANSLOCATING CHAIN-ASSOCIATING MEMBRANE PROTEIN).  
 9.5e-73:204:69  
 CANIS FAMILIARIS (DOG).  
 Q01685

25 F-HEMBB1000881  
 F-SPONDIN PRECURSOR.  
 1.2e-23:191:37  
 XENOPUS LAEVIS (AFRICAN CLAWED FROG).  
 P35447

30 F-HEMBB1000905  
 TRANSCRIPTIONAL REPRESSOR RCO-1.  
 0.068:105:34  
 NEUROSPORA CRASSA.  
 P78706

35 F-HEMBB1001026  
 ENDOSOMAL P24A PROTEIN PRECURSOR (70 KD ENDOMEMBRANE PROTEIN) (PHEROMONE ALPHA-  
 FACTOR TRANSPORTER) (ACIDIC 24 KD LATE ENDOCYTIC INTERMEDIATE COMPONENT).  
 1.3e-11:138:31  
 SACCHAROMYCES CEREVISIAE (BAKER'S YEAST).  
 P32802

40 F-HEMBB1001048  
 SARCALUMENIN PRECURSOR.  
 3.1e-20:151:32  
 ORYCTOLAGUS CUNICULUS (RABBIT).  
 P13666

45 F-HEMBB1001200  
 HYPOTHETICAL GENE 51 MEMBRANE PROTEIN.  
 1.0:66:27  
 55 ICTALURID HERPESVIRUS 1 (CHANNEL CATFISH VIRUS) (CCV).  
 Q00135

F-HEMBB1001407

F-Y79AA1002378  
ZINC FINGER PROTEIN 38 (ZFP-38) (CTFIN51) (TRANSCRIPTION FACTOR RU49).  
1.0e-59:163:74  
MUS MUSCULUS (MOUSE).  
Q07231

5

F-Y79AA1002381  
CELL DIVISION CONTROL PROTEIN 28 (EC 2.7.1.-).  
9.5e-41:179:38  
SACCHAROMYCES CEREVISIAE (BAKER'S YEAST).  
P00546

10

Homology search result 6

15 **[0296]** The result of the homology search in the GenBank(<http://www.ncbi.nlm.nih.gov/web/GenBank/>) using the clone sequences of the 5'-ends. except EST and STS sequences

Indicated are from the top,  
the name of the clone sequence,  
definition of the top hit data,  
the P-value: the length of the sequence used for comparison (nucleotide):similarity (%),  
the Accession No. of the top hit data.

20

**[0297]** Data were not shown for the clones in which the P-value was higher than 1.

25

F-BNGH41000020  
H.sapiens mitochondrial DNA, complete genome.  
6.0e-188:913:97  
X93334

30

F-BNGH41000087  
Human DNA sequence from clone 1049G16 on chromosome 20q12-13.2 Contains gene similar to GLU-COSAMINE-6-SULFATASE, a nuclear receptor coactivator gene, ESTs, STSs, GSSs, complete sequence.  
7.1e-32:176:99  
AL034418

35

F-BNGH41000091  
Homo sapiens potassium channel h-eag.  
1.6e-79:687:76  
AJ001366

40

F-HEMBA1000006  
S.erythraea second and third ORF's of eryA gene, complete cds.  
0.95:243:64  
M63677

45

F-HEMBA1000121  
Human DNA sequence \*\*\* SEQUENCING IN PROGRESS \*\*\* from clone 796F18, WORKING DRAFT SEQUENCE.  
5.9e-70:450:89  
AL031291

50

F-HEMBA1000128  
Plasmodium falciparum DNA \*\*\* SEQUENCING IN PROGRESS \*\*\* from contig 3-14, complete sequence.  
1.0:274:59  
Z98549

55

F-HEMBA1000275



- F-HEMBB1000106  
Human DNA sequence from clone 1170D6 on chromosome Xq22.3-23. Contains a pseudogene similar to U-SNRNP-associated Cyclophilin (USA-CYP, EC 5.2.1.8), ESTs, an STS and a GSS, complete sequence.  
0.033:332:61  
5 AL030995
- F-HEMBB1000276  
Dictyostelium discoideum gene encoding a novel glycoprotein.  
0.00070:440:60  
10 AJ005262
- F-HEMBB1000309  
Homo sapiens zinc finger protein (MBLL) mRNA, complete cds.  
7.6e-34:180:100  
15 AF061261
- F-HEMBB1000407  
Human Chromosome 11 pac pDJ393o15, WORKING DRAFT SEQUENCE, 8 unordered pieces. 0.16:228:64  
AC000384  
20
- F-HEMBB1000447  
Homo sapiens JWA protein mRNA, complete cds.  
1.4e-158:750:98  
25 AF070523
- F-HEMBB1000542  
Human DNA sequence from PAC 509L4 on chromosome 6q22.1-6q22.33. Contains SSX3 like pseudogene, EST, STS.  
4.3e-141:874:89  
30 Z99496
- F-HEMBB1000567  
Human DNA for insulin-like growth factor II (IGF-2); exon 7 and additional ORF.  
9.7e-122:572:99  
35 X07868
- F-HEMBB1000642
- F-HEMBB1000668  
Caenorhabditis elegans cosmid K06A5.  
0.00041:174:64  
40 AF039038
- F-HEMBB1000679  
C.familiaris mRNA for TRAM-protein.  
6.1e-100:756:80  
45 X63678
- F-HEMBB1000881  
Danio rerio mRNA for MINDIN2, complete cds.  
6.2e-40:581:66  
50 AB006085
- F-HEMBB1000905  
Homo sapiens clone RG315H11, WORKING DRAFT SEQUENCE, 5 unordered pieces.  
4.9e-91:209:94  
55 AC005089

1.1e-132:805:88  
U39045

5 F-Y79AA1002378  
Mus musculus mRNA for zinc finger protein, complete cds, clone:CTfin51.  
1.9e-64:521:78  
D10630

10 F-Y79AA1002381  
O.sativa mRNA for cdc2+/CDC28-related protein kinase.  
3.3e-21:431:60  
X58194

Homology search result 7

15 [0298] The result of the homology search in the GenBank(<http://www.ncbi.nlm.nih.gov/web/GenBank/>) using the clone sequences of the 3'-ends. except EST and STS sequences.

20 Indicated are from the top,  
the name of the clone sequence,  
definition of the top hit data,  
the P-value: the length of the sequence used for comparison (nucleotide):similarity (%),  
the Accession No. of the top hit data.

25 [0299] Blank indicates that the 3'-end sequence corresponding to the 5'-end was not determined in the clone. Data were not shown for the clones in which the P-value was higher than 1.

R-HEMBA1000006

30 R-HEMBA1000121  
Human DNA sequence \*\*\* SEQUENCING IN PROGRESS \*\*\* from clone 796F18, WORKING DRAFT SE-  
QUENCE.  
2.2e-43:355:80  
AL031291

35 R-HEMBA1000128  
Homo sapiens chromosome X, PAC 671D9, complete sequence.  
0.99:389:60  
AF031078

40 R-HEMBA1000275  
Homo sapiens DNA sequence from PAC 958B3 on chromosome Xp22.11-Xp22.22. Contains ESTs STS and CpG  
island.  
3.4e-10:212:66  
45 Z93023

R-HEMBA1000300  
[Alu RNA transcript, clone NE461] [human, embryonal carcinoma cells, NTera2D1 pluripotent cells, Other RNA;  
282 nt].  
50 4.6e-42:246:89  
S42653

R-nnnnnnnnnnnnn  
Homo sapiens chromosome 17, clone hRPK.235\_I\_10, complete sequence.  
55 1.0e-71:192:95  
AC005922

R-HEMBA1000462

- Homo sapiens JWA protein mRNA, complete cds.  
1.7e-107:533:97  
AF070523
- 5 R-HEMBB1000542  
Mus musculus bromodomain-containing protein BP75 mRNA, complete cds.  
4.4e-72:547:80  
AF084259
- 10 R-HEMBB1000567  
Human insulin-like growth factor (IGF-II) gene, exon 1 of 4.  
4.3e-60:368:88  
M13970
- 15 R-HEMBB1000642  
Human DNA sequence from PAC 46H23, BRCA2 gene region chromosome 13q12-13 contains Klotho, ESTs.  
2.9e-42:431:75  
Z84483
- 20 R-HEMBB1000668  
CITBI-E1-2508D15.TR CITBI-E1 Homo sapiens genomic clone 2508D15, genomic survey sequence.  
2.5e-40:249:91  
AQ261535
- 25 R-HEMBB1000679  
HS\_3061\_A1\_C03\_T7 CIT Approved Human Genomic Sperm Library D Homo sapiens genomic clone Plate=3061  
Col=5 Row=E, genomic survey sequence.  
1.8e-48:257:96  
AQ127602
- 30 R-HEMBB1000881  
CIT-HSP-2350020.TR CIT-HSP Homo sapiens genomic clone 2350020, genomic survey sequence.  
0.0072:248:61  
AQ062620
- 35 R-HEMBB1000905  
Homo sapiens clone RG315H11, WORKING DRAFT SEQUENCE, 5 unordered pieces.  
2.5e-104:547:94  
AC005089
- 40 R-HEMBB1001026  
  
R-HEMBB1001048
- 45 R-HEMBB1001200  
P.falciparum complete gene map of plastid-like DNA (IR-A).  
1.5e-11:521:59  
X95275
- 50 R-HEMBB1001407  
Human DNA sequence \*\*\* SEQUENCING IN PROGRESS \*\*\* from clone 2705, WORKING DRAFT SEQUENCE.  
3.0e-29:308:77  
AL033529
- 55 R-HEMBB1001530  
Homo sapiens chromosome 19, cosmid R30538, complete sequence.  
0.040:373:63  
AC005943

Homology search result 8.

**[0300]** The result of the homology search in the Human Unigene(<http://www.ncbi.nlm.nih.gov/UniGene>) using the clone sequences of the 5'-ends.

5

Indicated are from the top,  
the name of the clone sequence,  
title of the top hit data,  
the P-value: the length of the sequence used for comparison (nucleotide):similarity (%),  
the Accession No. of the top hit data.

10

**[0301]** Data were not shown for the clones in which the P-value was higher than 1.

15

F-BNGH41000020  
ESTs  
6.6e-72:412:92  
Hs.153375:AI287812

20

F-BNGH41000087  
Homo sapiens mRNA for MIFR-1, complete cds  
0.027:499:57  
Hs.58269:AB010962

25

F-BNGH41000091  
Homo sapiens voltage-gated potassium channel eag (EAG) mRNA, complete cds  
5.2e-81:687:76  
Hs.158305:AJ001366

30

F-HEMBA1000006  
ESTs, Weakly similar to HYPOTHETICAL 51.2 KD PROTEIN IN LAG1-RPL14B INTERGENIC REGION [S.cerevisiae]  
2.0e-25:167:91  
Hs.9252:R53360

35

F-HEMBA1000121  
ESTs, Moderately similar to HYPOTHETICAL 68.7 KD PROTEIN ZK757.1 IN CHROMOSOME III [Caenorhabditis elegans]  
3.0e-34:180:98  
Hs.149509:N24022

40

F-HEMBA1000128  
EST  
0.00069:177:62  
Hs.158854:AI377837

45

F-HEMBA1000275  
Human modulator recognition factor I (MRF-1) mRNA, 3'end  
0.012:508:58  
Hs.920:M62324

50

F-HEMBA1000300  
Human mRNA for KIAA0355 gene, complete cds  
1.6e-46:402:78  
Hs.153014:AB002353

55

F-HEMBA1000349  
EST  
6.7e-08:65:95

5 F-HEMBB1000567  
ESTs  
8.8e-13:271:71  
Hs.19934:AA455673

F-HEMBB1000642

10 F-HEMBB1000668  
EST  
0.83:192:58  
Hs.126372:AA912193

15 F-HEMBB1000679  
H.sapiens mRNA for TRAMP protein  
4.1e-96:727:80  
Hs.4147:X63679

20 F-HEMBB1000881  
Homo sapiens chromosome 4p homeobox mRNA sequence  
2.2e-06:512:60  
Hs.104134:M99587

25 F-HEMBB1000905  
Homo sapiens mRNA for voltage gated potassium channel  
0.93:337:58  
Hs.4975:Y15065

30 F-HEMBB1001026  
Human p76 mRNA, complete cds  
6.1e-08:410:61  
Hs.28757:U81006

35 F-HEMBB1001048  
Human Hpast (HPAST) mRNA, complete cds  
2.1e-56:524:75  
Hs.155119:AF001434

40 F-HEMBB 1001200  
EST  
0.10:300:61  
Hs.161647:AA133367

45 F-HEMBB1001407  
Homo sapiens PRKY exon 1 and joined CDS  
2.6e-40:271:81  
Hs.56336:Y15801

50 F-HEMBB1001530  
ESTs  
1.2e-98:477:98  
Hs.135208:AI093908

55 F-HEMBB1001547

F-HEMBB1001573  
EST  
2.2e-06:115:75  
Hs.138275:R43976

B94 PROTEIN  
5.7e-13:469:65  
Hs.75522:M92357

5 F-Y79AA1002058  
Homo sapiens clone 24733 mRNA sequence  
1.7e-154:740:98  
Hs.21970:AF052149

10 F-Y79AA1002121  
EST  
0.14:104:66  
Hs.100070:M91493

15 F-Y79AA1002129  
ESTs  
5.1e-90:431:98  
Hs.40719:AI183452

20 F-Y79AA1002213  
  
F-Y79AA1002334  
ESTs  
5.0e-20:187:80

25 Hs.111900:AA397579  
  
F-Y79AA1002373  
ESTs  
4.5e-37:192:98

30 Hs.118559:AA887084  
  
F-Y79AA1002376  
Homo sapiens cytoplasmic dynein intermediate chain 1 mRNA, complete cds  
1.2e-36:657:64

35 Hs.65248:AF063228  
  
F-Y79AA1002378  
Homo sapiens KIAA0426 mRNA, complete cds  
4.9e-38:424:72

40 Hs.97476:AB007886  
  
F-Y79AA1002381  
CELL DIVISION PROTEIN KINASE 3  
8.4e-17:580:61

45 Hs.100009:X66357

## Homology search result 9

50 [0302] The result of the homology search in the Human Unigene(<http://www.ncbi.nlm.nih.gov/UniGene>) using the clone sequences of the 3'-ends.

Indicated are from the top,  
the name of the clone sequence,  
title of the top hit data,  
55 the P-value: the length of the sequence used for comparison (nucleotide):similarity (%),  
the Accession No. of the top hit data.

[0303] Blank indicates that the 3'-end sequence corresponding to the 5'-end was not determined in the clone.

5 R-HEMBB1000567  
Insulin-like growth factor 2 (somatomedin A)  
8.9e-61:369:88  
Hs.155487:J03242

10 R-HEMBB1000642  
ESTs  
2.2e-44:308:84  
Hs.141318:N71080

15 R-HEMBB1000668  
ESTs, Weakly similar to hTAFII100 [H.sapiens]  
2.5e-102:520:95  
Hs.3830:AA167691

20 R-HEMBB1000679  
ESTs  
6.7e-36:188:97  
Hs.154218:AA169554

25 R-HEMBB1000881  
ESTs  
8.4e-105:519:96  
Hs.110967:AA570505

30 R-HEMBB1000905  
ESTs  
1.1e-94:454:98  
Hs.52515:AA464314

35 R-HEMBB1001026  
ESTs  
0.22:93:69  
Hs.119510:AA630235

40 R-HEMBB1001048  
EST  
0.42:127:66  
Hs.147466:AI215091

45 R-HEMBB1001200  
ESTs  
3.7e-07:330:62  
Hs.10109:AI148628

50 R-HEMBB1001407  
MHC class II transactivator  
3.8e-35:414:71  
Hs.3076:U18259

55 R-HEMBB1001530  
ESTs  
2.4e-95:455:98  
Hs.8956:AI146421

R-HEMBB1001547  
ESTs  
1.0e-111:533:98

# EP 1 130 094 A2

Homo sapiens DNA recombination and repair protein (MRE11B) mRNA, complete cds  
0.00075:456:59  
Hs.153855:AF022778

5 R-Y79AA1002213  
Human mRNA for KIAA0392 gene, partial cds  
6.2e-45:304:85  
Hs.40100:AB002390

10 R-Y79AA1002334  
ESTs  
7.7e-91:495:92  
Hs.90804:W28091

15 R-Y79AA1002373  
Human kpni repeat mrna (cdna clone pcd-kpni-8), 3' end  
5.2e-98:545:91  
Hs.103948:K00627

20 R-Y79AA1002376  
ESTs  
2.0e-91:455:97  
Hs.153375:AI287812

25 R-Y79AA1002378  
ESTs, Highly similar to ZINC FINGER PROTEIN ZFP-35 [Mus musculus]  
9.4e-15:131:83  
Hs.20082:W89121

30 R-Y79AA1002381  
ESTs, Highly similar to CELL DIVISION CONTROL PROTEIN 2 HOMOLOG [Plasmodium falciparum (isolate k1/thailand)]  
1.5e-104:531:95  
Hs.26322:AA156858

35 Homology search result 10

[0305] Data obtained by the homology search for full length nucleotide sequences and deduced amino acid sequences. In the result of the search shown below, both units, aa and bp, are used as length units for the sequences to be compared. Each data includes Clone name, Definition in matching data, P value, Length of sequence to be compared, Homology, and Accession number (No.) of matching data. These items are shown in this order, separated by a double-slash mark, //.

45 C-HEMBA1000006//Homo sapiens mRNA; cDNA DKFZp564G1762 (from clone DKFZp564G1762).//0//1230bp//92%//AB026894  
C-nnnnnnnnnnnnn//GAMETOGENESIS EXPRESSED PROTEIN GEG-154.//2.30E-71//344aa//50%//P50636  
C-HEMBA1000121//HYPOTHETICAL 68.7 KD PROTEIN ZK757.1 IN CHROMOSOME III.//4.80E-05//83aa//27%//P34679  
50 C-HEMBA1000128//PATHOGENESIS-RELATED PROTEIN 1 PRECURSOR (PR-1).//3.20E-07//89aa//34%//P33154  
C-HEMBA1000275  
C-HEMBA1000300  
C-HEMBA1000349//ATP-BINDING CASSETTE TRANSPORTER 1.//5.30E-65//352aa//39%//P41233  
C-HEMBA1000443//Homo sapiens CGI-96 protein mRNA, complete cds.//4.70E-129//686bp//91%//AF151854  
55 C-HEMBA1000590//Homo sapiens mRNA for matrilin-4, partial.//2.00E-273//1254bp//99%//AJ007581  
C-HEMBA1000634//Homo sapiens T-cell activation protein (PGR1) gene, complete cds.//0//994bp//99%//AF116272  
C-HEMBA1000713//Homo sapiens 10kD protein (BC10) mRNA, complete cds.//0//1254bp//99%//AF053470



C-HEMBA1000745//COLLAGEN ALPHA 1(I) CHAIN (FRAGMENTS).//2.00E-07//445aa//27%/P02454  
 C-HEMBA1000907  
 C-HEMBA1000940//GAP JUNCTION ALPHA-3 PROTEIN (CONNEXIN 44) (CX44).//2.90E-39//362aa//31%/P41987  
 5 C-HEMBA1000962  
 C-HEMBA1001221//AGRN PRECURSOR.//2.50E-25//294aa//29%/P31696  
 C-HEMBA1001228//Human germline oligomeric matrix protein (COMP) mRNA, complete cds.//7.80E-286//1105bp//94%/L32137  
 C-HEMBA1001297  
 10 C-HEMBA1001390//Mus musculus polymerase I-transcript release factor mRNA, complete cds.//2.50E-57//464bp//82%/AF036249  
 C-HEMBA1001563  
 C-HEMBA1001621//PROBABLE G PROTEIN-COUPLED RECEPTOR APJ.//3.50E-123//259aa//89%/P35414  
 C-nnnnnnnnnnn//ZINC FINGER PROTEIN 84 (ZINC FINGER PROTEIN HPF2).//2.40E-85//293aa//50%/P51523  
 15 C-HEMBA1001878//Homo sapiens U5 snRNP-specific 40 kDa protein mRNA, complete cds.//0//1488bp//99%/AF090988  
 C-HEMBA1002131//PROCOLLAGEN-LYSINE,2-OXOGLUTARATE 5-DIOXYGENASE 1//4.10E-10//140aa//30%/P24802  
 20 C-HEMBA1002163//HYPOTHETICAL 40.7 KD PROTEIN IN DAK1-ORC1 INTERGENIC REGION.//9.40E-28//309aa//30%/Q04651  
 C-HEMBA1002164  
 C-HEMBA1002167//Rattus norvegicus neuroligin I mRNA, complete cds.//1.30E-305//1643bp//91%/U22952  
 C-HEMBA1002178//PROCOLLAGEN-LYSINE,2-OXOGLUTARATE 5-DIOXYGENASE 1 PRECURSOR (EC 1.14.11.4) (LYSYL HYDROXYLASE 1) (LH1).//3.70E-10//140aa//30%/P24802  
 25 C-nnnnnnnnnnn//Human glycyl-tRNA synthetase mRNA, complete cds.//0//2380bp//99%/U09587  
 C-HEMBA1002195//VEGETATIBLE INCOMPATIBILITY PROTEIN HET-E-1.//8.80E-23//221aa//31%/Q00808  
 C-HEMBA1002227//Homo sapiens mRNA for 80K-L protein, complete cds.//0//1324bp//98%/D10522  
 C-HEMBA1002239  
 30 C-HEMBA1002316  
 C-HEMBA1002420  
 C-HEMBA1002421//Human heparan sulfate proteoglycan (HSPG) core protein, 3' end.//0//2097bp//99%/J04621  
 C-HEMBA1002524//Human MHC Class I region proline rich protein mRNA, complete cds.//0//1763bp//95%/U63336  
 35 C-HEMBA1002551//PUTATIVE SERINE/THREONINE-PROTEIN KINASE PKWA (EC 2.7.1.-).//9.80E-08//110aa//37%/P49695  
 C-HEMBA1002767//Homo sapiens chromosome 1p33-p34 beta-1,4-galactosyltransferase mRNA, complete cds.//0//1497bp//99%/AF038660  
 C-HEMBA1002992//UBIQUITIN-LIKE PROTEIN DSK2.//2.00E-21//216aa//35%/P48510  
 40 C-HEMBA1003047//Homo sapiens intrinsic factor-B12 receptor precursor, mRNA, complete cds.//0//1768bp//99%/AF034611  
 C-HEMBA1003072//Gallus gallus single-strand DNA-binding protein csdp mRNA, partial cds.//3.30E-93//927bp//73%/U68380  
 C-HEMBA1003101//Homo sapiens tyrosylprotein sulfotransferase-2 mRNA, complete cds.//0//1854bp//99%/AF049891  
 45 C-HEMBA1003230//Homo sapiens fibulin-5.//5.60E-308//1398bp//99%/AJ133490  
 C-HEMBA1003294  
 C-HEMBA1003315//Mus musculus mRNA for DNA helicase, complete cds.//6.30E-250//1426bp//88%/AB013912  
 C-HEMBA1003392//Homo sapiens LDL receptor-related protein 6 (LRP6) mRNA, complete cds.//0//1721bp//100%/AF074264  
 50 C-HEMBA1003399//MVP1 PROTEIN.//2.30E-15//279aa//23%/P40959  
 C-HEMBA1003487  
 C-HEMBA1003530//S.scrofa mRNA for BM88 antigen.//1.20E-60//900bp//66%/X82027  
 C-HEMBA1003602//Homo sapiens CGI-67 protein mRNA, complete cds.//3.50E-70//732bp//66%/AF151825  
 55 C-HEMBA1003732//SFT2 PROTEIN.//1.50E-06//162aa//30%/P38166  
 C-HEMBA1003945//Homo sapiens hypothetical 43.2 Kd protein mRNA, complete cds.//8.90E-287//757bp//97%/AF077030  
 C-HEMBA1004110//Homo sapiens intersectin short form mRNA, complete cds.//0//2033bp//99%/AF064243

C-HEMBA1004250//CADHERIN-RELATED TUMOR SUPPRESSOR PRECURSOR (FAT PROTEIN).//6.40E-51//  
277aa//35%/P33450  
C-HEMBA1004391//TUMOR SUPPRESSOR PROTEIN DCC PRECURSOR.//5.60E-20//194aa//26%/P70211  
C-HEMBA1004444//GLYCOPROTEIN 25L PRECURSOR (GP25L).//4.60E-41//148aa//52%/P27869  
5 C-HEMBA1004454  
C-HEMBA1004505//MANNOSYL-OLIGOSACCHARIDE ALPHA-1,2-MANNOSIDASE ISOFORM 1 (EC  
3.2.1.113) (MAN(9)-ALPHA-MANNOSIDASE).//2.70E-45//239aa//43%/P53624  
C-HEMBA1004797  
C-HEMBA1004982//TETRACYCLINE RESISTANCE PROTEIN, CLASS E (TETA(E)).//6.30E-10//149aa//26%/  
10 Q07282  
C-HEMBA1005070//SKIN SECRETORY PROTEIN XP2 PRECURSOR (APEG PROTEIN).//1.10E-05//187aa//  
29%/P17437  
C-HEMBA1005084//NON-RECEPTOR TYROSINE KINASE SPORE LYSIS A (EC 2.7.1.112) (TYROSINE- PRO-  
TEIN KINASE 1).//1.20E-07//102aa//37%/P18160  
15 C-HEMBA1005145  
C-HEMBA1005430  
C-HEMBA1005449//GLUCOAMYLASE S1/S2 PRECURSOR (EC 3.2.1.3) (GLUCAN 1,4-ALPHA- GLUCOSI-  
DASE) (1,4-ALPHA-D-GLUCAN GLUCOHYDROLASE).//5.40E-10//224aa//24%/P13983  
C-HEMBA1005489  
20 C-HEMBA1005522//COAGULATION FACTOR VII PRECURSOR (EC 3.4.21.21).//7.70E-15//78aa//51%/P98139  
C-HEMBA1005545//MUSCARINIC ACETYLCHOLINE RECEPTOR M3.//0//2121aa//100%/U29589  
C-HEMBA1005698//Homo sapiens vesicle trafficking protein (SEC22C) mRNA, complete cds.//6.60E-163//  
753bp//99%/AF039568  
C-HEMBA1005913  
25 C-HEMBA1005929//H.sapiens mRNA for serine/threonine protein kinase EMK.//6.50E-92//1092bp//69%/X97630  
C-HEMBA1005945//Oryctolagus cuniculus peroxisomal Ca-dependent solute carrier mRNA, complete Gds.//  
1.90E-44//666bp//65%/AF004161  
C-HEMBA1006016  
C-HEMBA1006171  
30 C-HEMBA1006299  
C-HEMBA1006311  
C-HEMBA1006335  
C-HEMBA1006430//Human putative transmembrane protein precursor (B5) mRNA, complete cds.//2.40E-70//  
1108bp//65%/L38961  
35 C-HEMBA1006482//Homo sapiens h-scol (SCOI) mRNA, nuclear gene encoding mitochondrial protein, complete  
cds.//0//1101bp//98%/AF026852  
C-HEMBA1006572//ODD-SKIPPED PROTEIN.//2.60E-39//85aa//83%/P23803  
C-HEMBA1006707//Homo sapiens mRNA for matrilin-4, partial.//0//2003bp//99%/AJ007581  
C-HEMBA1006724  
40 C-HEMBA1006902//Homo sapiens mRNA for matrilin-4, partial.//4.80E-275//1799bp//85%/AJ007581  
C-HEMBA1006916//Homo sapiens Grb14 mRNA, complete cds.//3.00E-277//1010bp//95%/L76687  
C-HEMBA1006960  
C-HEMBA1007013  
C-HEMBA1007057  
45 C-HEMBA1007241  
C-HEMBA1007291  
C-HEMBA1007332  
C-HEMBB1000276  
C-HEMBB1000447//Homo sapiens JWA protein mRNA, complete cds.//0//2059bp//99%/AF070523  
50 C-HEMBB1000642  
C-HEMBB1000668//Homo sapiens mRNA for KIAA0893 protein, complete cds.//0//2375bp//99%/AB020700  
C-HEMBB1000679//C.familiaris mRNA for TRAM-protein.//4.10E-210//1149bp//80%/X63678  
C-HEMBB1000881//Danio rerio mRNA for MINDIN2, complete cds.//1.70E-67//948bp//66%/AB006085  
C-HEMBB1000905//TRANSCRIPTIONAL REPRESSOR RCO-1.//1.00E-11//311aa//27%/P78706  
55 C-HEMBB1001026//ENDOSOMAL P24A PROTEIN PRECURSOR (70 KD ENDOMEMBRANE PROTEIN) (PHE-  
ROMONE ALPHA-FACTOR TRANSPORTER) (ACIDIC 24 KD LATE ENDOCYTIC INTERMEDIATE COMPO-  
NENT).//5.30E-11//142aa//30%/P32802  
C-HEMBB1001048//Human Hpast (HPAST) mRNA, complete cds.//6.50E-39//448bp//75%/AC000159

O15127  
 C-HEMBA1006430//Human putative transmembrane protein precursor (B5) mRNA, complete cds.//2.40E-70//1108bp//65%//L38961  
 C-HEMBA1006482//Homo sapiens h-sco1 (SCO1) mRNA, nuclear gene encoding mitochondrial protein, complete cds.//0//1101bp//98%//AF026852  
 5 C-HEMBA1006517  
 C-HEMBA1006544  
 C-HEMBA1006572//ODD-SKIPPED PROTEIN.//2.60E-39//85aa//83%//P23803  
 C-HEMBA1006658//Homo sapiens mRNA for NIK, partial cds.//0//1500bp//98%//AB013385  
 10 C-HEMBA1006707//Homo sapiens mRNA for matrilin-4, partial.//0//2003bp//99%//AJ007581  
 C-HEMBA1006724  
 C-HEMBA1006749//Homo sapiens mRNA for matrilin-4, partial.//1.40E-275//1942bp//83%//AJ007581  
 C-HEMBA1006770//FLOWERING TIME CONTROL PROTEIN FCA.//1.20E-33//352aa//34%//O04425  
 C-HEMBA1006902//Homo sapiens mRNA for matrilin-4, partial.//4.80E-275//1799bp//85%//AJ007581  
 15 C-HEMBA1006912  
 C-HEMBA1006916//Homo sapiens Grb14 mRNA, complete cds.//3.00E-277//1010bp//95%//L76687  
 C-HEMBA1006960  
 C-HEMBA1007013//Mus musculus sphingosine kinase (SPHK1b) mRNA, complete cds.//1.10E-14//412bp//63%//AF068749  
 20 C-HEMBA1007057  
 C-HEMBA1007063  
 C-HEMBA1007226//Homo sapiens RPA-binding trans-activator (RBT1) mRNA, complete cds.//7.30E-273//1242bp//99%//AF192529  
 C-HEMBA1007241//HYPOTHETICAL 24.5 KD PROTEIN IN SAP185-BCK1 INTERGENIC REGION.//2.70E-14//106aa//42%//P40857  
 25 C-HEMBA1007291  
 C-HEMBA1007332//Homo sapiens mRNA for unr-interacting protein.//6.40E-83//266bp//98%//AJ010025  
 C-HEMBA1000106//CELLULAR NUCLEIC ACID BINDING PROTEIN (CNBP).//1.60E-10//139aa//30%//P53996  
 C-HEMBA1000276  
 30 C-HEMBA1000309  
 C-HEMBA1000407  
 C-HEMBA1000447//Homo sapiens JWA protein mRNA, complete cds.//0//2059bp//99%//AF070523  
 C-HEMBA1000542//Mus musculus bromodomain-containing protein BP75 mRNA, complete cds.//2.60E-232//1452bp//85%//AF084259  
 35 C-HEMBA1000567  
 C-HEMBA1000642  
 C-HEMBA1000668//Homo sapiens mRNA for KIAA0893 protein, complete cds.//0//2375bp//99%//AB020700  
 C-HEMBA1000679//C.familiaris mRNA for TRAM-protein.//4.10E-210//1149bp//80%//X63678  
 C-HEMBA1000881//Danio rerio mRNA for MINDIN2, complete cds.//1.70E-67//948bp//66%//AB006085  
 40 C-HEMBA1000905//TRANSCRIPTIONAL REPRESSOR RCO-1.//1.00E-11//311aa//27%//P78706  
 C-HEMBA1001026//ENDOSOMAL P24A PROTEIN PRECURSOR (70 KD ENOMEMBRANE PROTEIN) (PHE-  
 ROMONE ALPHA-FACTOR TRANSPORTER) (ACIDIC 24 KD LATE ENDOCYTIC INTERMEDIATE COMPO-  
 NENT).//5.30E-11//142aa//30%//P32802  
 C-HEMBA1001048//SARCALUMENIN PRECURSOR.//6.50E-18//154aa//33%//P13666  
 45 C-HEMBA1001200  
 C-HEMBA1001407  
 C-HEMBA1001530//SLS1 PROTEIN PRECURSOR.//9.80E-10//273aa//27%//Q99158  
 C-HEMBA1001547//Homo sapiens CGI-02 protein mRNA, complete cds.//0//2311bp//99%//AF132937  
 C-HEMBA1001573  
 50 C-HEMBA1001847//NEUROGENIC PROTEIN BIG BRAIN.//4.70E-06//258aa//24%//P23645  
 C-HEMBA1001959//CCAAT-BINDING TRANSCRIPTION FACTOR SUBUNIT A (CBF-A) (NF-Y PROTEIN CHAIN  
 B) (NF-YB) (CAAT-BOX DNA BINDING PROTEIN SUBUNIT B).//7.30E-14//97aa//38%//P25210  
 C-HEMBA1001978  
 C-HEMBA1002039  
 55 C-HEMBA1002041//Homo sapiens transmembrane protein TENB2 (TENB2) mRNA, complete cds.//0//1746bp//99%//AF179274  
 C-HEMBA1002051//Homo sapiens Ets transcription factor ESE-2b mRNA, complete cds.//1.30E-95//454bp//99%//AF115403

NASE I).//1.00E-77//359aa//44%//Q14012  
 C-Y79AA1001013  
 C-Y79AA1001056//Homo sapiens MAID protein mRNA, complete cds.//0//1475bp//99%//AF113535  
 C-Y79AA1001062//TUMOR NECROSIS FACTOR, ALPHA-INDUCED PROTEIN 1, ENDOTHELIAL (B12 PRO-  
 5 TEIN).//8.90E-12//132aa//38%//Q13829  
 C-Y79AA1001090//NUCLEAR FACTOR NF-KAPPA-B P105 SUBUNIT (DNA-BINDING FACTOR KBF1) (EBP- 1)  
 (NF-KAPPA-B1 P84/NF-KAPPA-B1 P98) [CONTAINS: NUCLEAR FACTOR NF- KAPPA-B P50 SUBUNIT] (FRAG-  
 MENT).//4.50E-09//144aa//31%//Q63369  
 C-Y79AA1001212//Homo sapiens SL15 protein mRNA, complete cds.//6.30E-306//1388bp//99%//AF038961  
 10 C-Y79AA1001264//HYPOTHETICAL 39.9 KD PROTEIN T15H9.1 IN CHROMOSOME II PRECURSOR.//5.10E-  
 106//351aa//58%//Q10005  
 C-Y79AA1001272//Homo sapiens retinoic acid repressible protein (RARG-1) mRNA, complete cds.//1.50E-183//  
 867bp//98%//AF172066  
 C-Y79AA1001328//Mus musculus mRNA for Dll3 protein, complete cds.//1.90E-263//1988bp//79%//AB013440  
 15 C-Y79AA1001426//ANION EXCHANGE PROTEIN 3 (CARDIAC/BRAIN BAND 3-LIKE PROTEIN) (CAE3/BAE3).//  
 6.20E-66//609aa//31%//P48751  
 C-Y79AA1001427//Homo sapiens cytochrome b5 reductase 1 (B5R.1) mRNA, complete cds.//0//1588bp//99%//  
 AF169481  
 C-Y79AA1001430//Homo sapiens mRNA for KIAA0469 protein, complete cds.//0//2943bp//99%//AB007938  
 20 C-Y79AA1001523//Homo sapiens transcriptional intermediary factor 1 alpha mRNA, complete cds.//0//2263bp//  
 99%//AF119042  
 C-Y79AA1001530//Human beta-tubulin gene (5-beta) with ten Alu family members.//0//1920bp//98%//X00734  
 C-Y79AA1001592  
 C-Y79AA1001727//CELL SURFACE A33 ANTIGEN PRECURSOR.//1.10E-13//286aa//27%//Q99795  
 25 C-Y79AA1001787//PROBABLE CALCIUM-TRANSPORTING ATPASE 9 (EC 3.6.1.38).//1.70E-133//544aa//37%//  
 Q12697  
 C-Y79AA1001793//Mus musculus mRNA for GSG1, complete cds.//3.70E-126//532bp//78%//D87325  
 C-Y79AA1001795//Homo sapiens mRNA for GalT4 protein.//2.30E-250//1137bp//99%//Y15061  
 C-Y79AA1001799//MITOCHONDRIAL RNA SPLICING PROTEIN MSR4.//3.40E-54//182aa//39%//P23500  
 30 C-Y79AA1001803//Homo sapiens secretogranin III mRNA, complete cds.//0//1871bp//99%//AF078851  
 C-Y79AA1001863  
 C-Y79AA1002022//POLIOVIRUS RECEPTOR HOMOLOG PRECURSOR.//2.20E-06//140aa//26%//P32507  
 C-Y79AA1002058//Mus musculus Gng3lg mRNA, complete cds.//4.10E-167//1145bp//83%//AF069954  
 C-Y79AA1002121//HISTONE H1.//4.90E-12//114aa//35%//P35060  
 35 C-Y79AA1002129  
 C-Y79AA1002213//HYPOTHETICAL 52.7 KD PROTEIN C38C10.2 IN CHROMOSOME III.//1.20E-98//262aa//  
 41%//Q03567  
 C-Y79AA1002334//GLUCOSE REPRESSION MEDIATOR PROTEIN.//1.70E-10//333aa//23%//P14922  
 C-Y79AA1002373//Mus musculus mRNA for GSG1, complete cds.//7.20E-147//680bp//79%//D87325  
 40 C-Y79AA1002376//Rattus norvegicus cytoplasmic dynein intermediate chain 2B mRNA, complete cds.//1.50E-  
 304//1667bp//90%//U39045  
 C-Y79AA1002378//Homo sapiens zinc finger protein NY-REN-21 antigen mRNA, partial cds.//0//963bp//99%//  
 AF155100  
 C-Y79AA1002381//Homo sapiens cell cycle related kinase mRNA, complete cds.//0//1791bp//98%//AF035013  
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## Claims

1. Use of an oligonucleotide as a primer for synthesizing the polynucleotide comprising the nucleotide sequence set forth in any one of SEQ ID NOs: 1-829 and 2545, or the complementary strand thereof, wherein said oligonucleotide is complementary to said polynucleotide or the complementary strand thereof and comprises at least 15 nucleotides.
2. A primer set for synthesizing polynucleotides, the primer set comprising an oligo-dT primer and an oligonucleotide complementary to the complementary strand of the polynucleotide comprising the nucleotide sequence set forth in any one of SEQ ID NOs: 1-829 and 2545, wherein said oligonucleotide comprises at least 15 nucleotides.
3. A primer set for synthesizing polynucleotides, the primer set comprising a combination of an oligonucleotide com-

prising a nucleotide sequence complementary to the complementary strand of the polynucleotide comprising a 5'-end nucleotide sequence and an oligonucleotide comprising a nucleotide sequence complementary to the polynucleotide comprising a 3'-end nucleotide sequence, wherein said oligonucleotides comprise at least 15 nucleotides and wherein said combination of 5'-end nucleotide sequence.3'-end nucleotide sequence is selected from the group consisting of:

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SEQ ID NO:4 and SEQ ID NO:830  
SEQ ID NO:5 and SEQ ID NO:831  
SEQ ID NO:6 and SEQ ID NO:832  
SEQ ID NO:7 and SEQ ID NO:833  
SEQ ID NO:8 and SEQ ID NO:834  
SEQ ID NO:9 and SEQ ID NO:835  
SEQ ID NO:11 and SEQ ID NO:836  
SEQ ID NO:12 and SEQ ID NO:837  
SEQ ID NO:13 and SEQ ID NO:838  
SEQ ID NO:14 and SEQ ID NO:839  
SEQ ID NO:15 and SEQ ID NO:840  
SEQ ID NO:16 and SEQ ID NO:841  
SEQ ID NO:17 and SEQ ID NO:842  
SEQ ID NO:18 and SEQ ID NO:843  
SEQ ID NO:20 and SEQ ID NO:844  
SEQ ID NO:22 and SEQ ID NO:845  
SEQ ID NO:23 and SEQ ID NO:846  
SEQ ID NO:24 and SEQ ID NO:847  
SEQ ID NO:25 and SEQ ID NO:848  
SEQ ID NO:26 and SEQ ID NO:849  
SEQ ID NO:27 and SEQ ID NO:850  
SEQ ID NO:28 and SEQ ID NO:851  
SEQ ID NO:29 and SEQ ID NO:852  
SEQ ID NO:30 and SEQ ID NO:853  
SEQ ID NO:31 and SEQ ID NO:854  
SEQ ID NO:32 and SEQ ID NO:855  
SEQ ID NO:33 and SEQ ID NO:856  
SEQ ID NO:34 and SEQ ID NO:857  
SEQ ID NO:35 and SEQ ID NO:858  
SEQ ID NO:36 and SEQ ID NO:859  
SEQ ID NO:37 and SEQ ID NO:860  
SEQ ID NO:39 and SEQ ID NO:861  
SEQ ID NO:40 and SEQ ID NO:862  
SEQ ID NO:41 and SEQ ID NO:863  
SEQ ID NO:42 and SEQ ID NO:864  
SEQ ID NO:44 and SEQ ID NO:865  
SEQ ID NO:45 and SEQ ID NO:866  
SEQ ID NO:46 and SEQ ID NO:867  
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SEQ ID NO:52 and SEQ ID NO:872  
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SEQ ID NO:54 and SEQ ID NO:874  
SEQ ID NO:55 and SEQ ID NO:875  
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SEQ ID NO:57 and SEQ ID NO:877  
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SEQ ID NO:60 and SEQ ID NO:880  
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SEQ ID NO:62 and SEQ ID NO:882  
SEQ ID NO:63 and SEQ ID NO:883  
SEQ ID NO:64 and SEQ ID NO:884  
SEQ ID NO:65 and SEQ ID NO:885  
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SEQ ID NO:67 and SEQ ID NO:887  
SEQ ID NO:69 and SEQ ID NO:888  
SEQ ID NO:70 and SEQ ID NO:889  
SEQ ID NO:71 and SEQ ID NO:890  
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SEQ ID NO:73 and SEQ ID NO:892  
SEQ ID NO:74 and SEQ ID NO:893  
SEQ ID NO:75 and SEQ ID NO:894  
SEQ ID NO:76 and SEQ ID NO:895  
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SEQ ID NO:79 and SEQ ID NO:898  
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SEQ ID NO:82 and SEQ ID NO:901  
SEQ ID NO:83 and SEQ ID NO:902  
SEQ ID NO:84 and SEQ ID NO:903  
SEQ ID NO:85 and SEQ ID NO:904  
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SEQ ID NO:87 and SEQ ID NO:906  
SEQ ID NO:88 and SEQ ID NO:907  
SEQ ID NO:89 and SEQ ID NO:908  
SEQ ID NO:90 and SEQ ID NO:909  
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SEQ ID NO:92 and SEQ ID NO:911  
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SEQ ID NO:94 and SEQ ID NO:913  
SEQ ID NO:95 and SEQ ID NO:914  
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SEQ ID NO:135 and SEQ ID NO:953  
SEQ ID NO:136 and SEQ ID NO:954  
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SEQ ID NO:138 and SEQ ID NO:956  
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10 SEQ ID NO:162 and SEQ ID NO:980  
SEQ ID NO:163 and SEQ ID NO:981  
SEQ ID NO:164 and SEQ ID NO:982  
SEQ ID NO:165 and SEQ ID NO:983  
SEQ ID NO:166 and SEQ ID NO:984  
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SEQ ID NO:168 and SEQ ID NO:986  
SEQ ID NO:169 and SEQ ID NO:987  
SEQ ID NO:170 and SEQ ID NO:988  
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20 SEQ ID NO:172 and SEQ ID NO:990  
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SEQ ID NO:178 and SEQ ID NO:996  
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SEQ ID NO:184 and SEQ ID NO:1002  
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SEQ ID NO:189 and SEQ ID NO:1007  
SEQ ID NO:190 and SEQ ID NO:1008  
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SEQ ID NO:194 and SEQ ID NO:1011  
SEQ ID NO:195 and SEQ ID NO:1012  
SEQ ID NO:196 and SEQ ID NO:1013  
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SEQ ID NO:203 and SEQ ID NO:1020  
SEQ ID NO:204 and SEQ ID NO:1021  
SEQ ID NO:205 and SEQ ID NO:1022  
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SEQ ID NO:208 and SEQ ID NO:1024



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 SEQ ID NO:827 and SEQ ID NO:1570  
 SEQ ID NO:828 and SEQ ID NO:1571  
 35 SEQ ID NO:829 and SEQ ID NO:1572, and  
 SEQ ID NO:2545 and SEQ ID NO:2546

4. A polynucleotide which can be synthesized with the primer set of claim 2 or 3.
5. A polynucleotide comprising a coding region in the polynucleotide of claim 4.
6. A substantially pure protein encoded by polynucleotide of claim 4.
7. A partial peptide of the protein of claim 6.
8. An isolated polynucleotide selected from the group consisting of
  - (a) a polynucleotide comprising a coding region of the nucleotide sequence set forth in any one of the following SEQ ID NOs:

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(b) a polynucleotide comprising a nucleotide sequence encoding a protein comprising the amino acid sequence

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and SEQ ID NO:4179

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(c) a polynucleotide comprising a nucleotide sequence encoding a protein comprising an amino acid sequence selected from the amino acid sequences of (b), in which one or more amino acids are substituted, deleted, inserted, and/or added, wherein said protein is functionally equivalent to the protein comprising said amino acid sequence selected from the amino acid sequences of (b);

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(d) a polynucleotide that hybridizes with a polynucleotide comprising a nucleotide sequence selected from the nucleotide sequences of (a), and that comprises a nucleotide sequence encoding a protein functionally equivalent to the protein encoded by the nucleotide sequence selected from the nucleotide sequences of (a);

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(e) a polynucleotide comprising a nucleotide sequence encoding a partial amino acid sequence of a protein encoded by the polynucleotide of (a) to (d);

(f) a polynucleotide comprising a nucleotide sequence with at least 70% identity to the nucleotide sequence of (a).

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9. A substantially pure protein encoded by the polynucleotide of claim 8.

10. An antibody against the protein or peptide of any one of claims 6, 7, and 9.

11. A vector comprising the polynucleotide of claim 5 or 8.

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12. A transformant carrying the polynucleotide of claim 5 or 8, or the vector of claim 11.

13. A transformant expressively carrying the polynucleotide of claim 5 or 8, or the vector of claim 11.

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14. A method for producing the protein or peptide of any one of claims 6, 7, and 9, comprising culturing the transformant of claim 13 and recovering the expression product.

15. An oligonucleotide comprising the nucleotide sequence of claim 8 (a) or the nucleotide sequence complementary to the complementary strand thereof, wherein said oligonucleotide comprises 15 nucleotides or more.

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16. Use of the oligonucleotide of claim 15 as a primer for synthesizing a polynucleotide.

17. Use of the oligonucleotide of claim 15 as a probe for detecting a gene.

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18. An antisense polynucleotide against the polynucleotide of claim 8, or the portion thereof.

19. A method for synthesizing a polynucleotide, the method comprising:

a) synthesizing a complementary strand using a cDNA library as a template, and using the primer set of claim 2 or 3, or the primer of claim 16; and

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b) recovering the synthesized product.

20. The method of claim 19, wherein the cDNA library is obtainable by oligo-capping method.

21. The method of claim 19, wherein the complementary strand is obtainable by PCR.

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22. A method for detecting the polynucleotide of claim 8, the method comprising:

a) incubating a target polynucleotide with the oligonucleotide of claim 15 under the conditions where hybridization occurs, and

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b) detecting the hybridization of the target polynucleotide with the oligonucleotide of claim 15.

23. A database of polynucleotides and/or proteins, the database comprising information on at least one sequence selected from the nucleotide sequences of claim 8 (a) and/or the amino acid sequences of claim 8 (b), or a medium

on which the database is stored.

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Figure 1

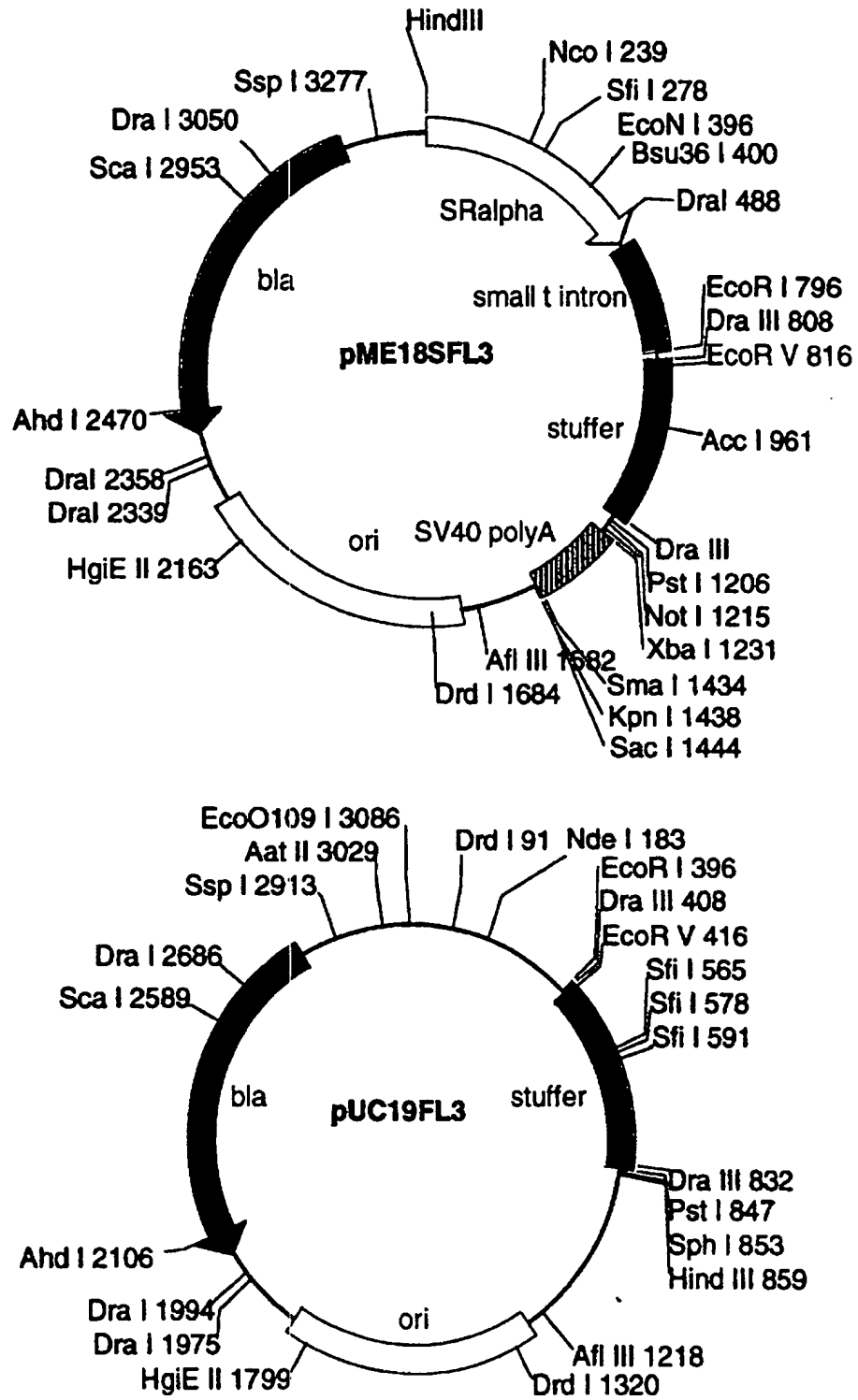


Figure 2

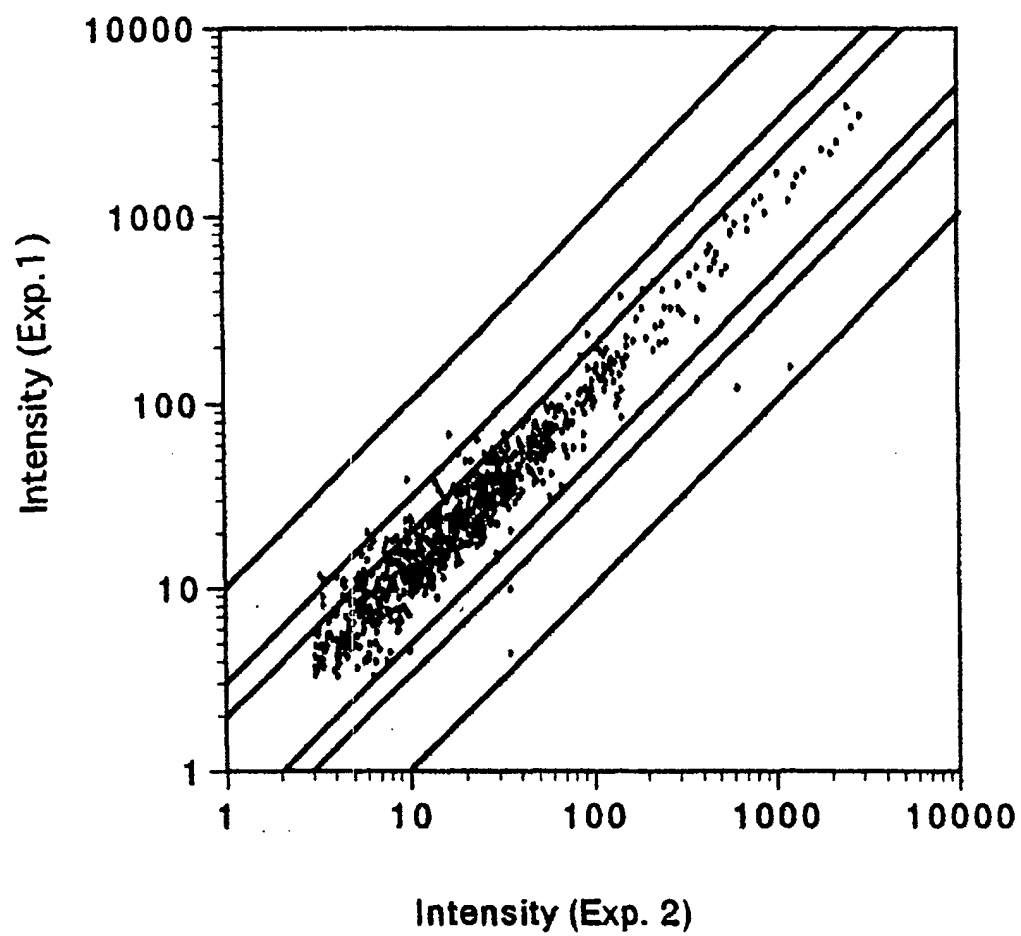


Figure 3

